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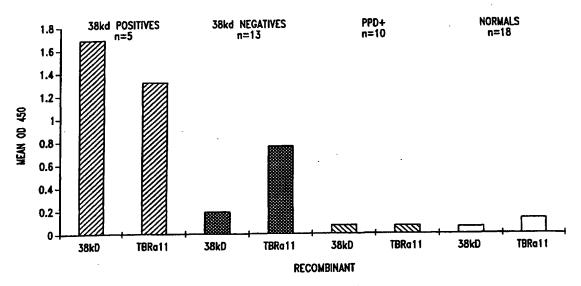
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(54) Title: COMPOUNDS AND METHODS FOR DIAGNOSIS OF TUBERCULOSIS



(57) Abstract

Compounds and methods for diagnosing tuberculosis are disclosed. The compounds provided include polypeptides that contain at least one antigenic portion of one or more M. tuberculosis proteins, and DNA sequences encoding such polypeptides. Diagnostic kits containing such polypeptides or DNA sequences and a suitable detection reagent may be used for the detection of M. tuberculosis infection in patients and biological samples. Antibodies directed against such polypeptides are also provided.

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COMPOUNDS AND METHODS FOR DIAGNOSIS OF TUBERCULOSIS

TECHNICAL FIELD

The present invention relates generally to the detection of *Mycobacterium* tuberculosis infection. The invention is more particularly related to polypeptides comprising a *Mycobacterium tuberculosis* antigen, or a portion or other variant thereof, and the use of such polypeptides for the serodiagnosis of *Mycobacterium tuberculosis* infection.

BACKGROUND OF THE INVENTION

Tuberculosis is a chronic, infectious disease, that is generally caused by infection with *Mycobacterium tuberculosis*. It is a major disease in developing countries, as well as an increasing problem in developed areas of the world, with about 8 million new cases and 3 million deaths each year. Although the infection may be asymptomatic for a considerable period of time, the disease is most commonly manifested as an acute inflammation of the lungs, resulting in fever and a nonproductive cough. If left untreated, serious complications and death typically result.

Although tuberculosis can generally be controlled using extended antibiotic therapy, such treatment is not sufficient to prevent the spread of the disease. Infected individuals may be asymptomatic, but contagious, for some time. In addition, although compliance with the treatment regimen is critical, patient behavior is difficult to monitor. Some patients do not complete the course of treatment, which can lead to ineffective treatment and the development of drug resistance.

Inhibiting the spread of tuberculosis will require effective vaccination and accurate, early diagnosis of the disease. Currently, vaccination with live bacteria is the most efficient method for inducing protective immunity. The most common Mycobacterium for this purpose is Bacillus Calmette-Guerin (BCG), an avirulent strain of *Mycobacterium bovis*. However, the safety and efficacy of BCG is a source of controversy and some countries, such as the United States, do not vaccinate the general public. Diagnosis is commonly achieved using a skin test, which involves intradermal exposure to tuberculin PPD (protein-purified derivative). Antigen-specific T cell responses result in measurable incubation at the injection

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site by 48-72 hours after injection, which indicates exposure to Mycobacterial antigens. Sensitivity and specificity have, however, been a problem with this test, and individuals vaccinated with BCG cannot be distinguished from infected individuals.

While macrophages have been shown to act as the principal effectors of *M. tuberculosis* immunity, T cells are the predominant inducers of such immunity. The essential role of T cells in protection against *M. tuberculosis* infection is illustrated by the frequent occurrence of *M. tuberculosis* in AIDS patients, due to the depletion of CD4 T cells associated with human immunodeficiency virus (HIV) infection. Mycobacterium-reactive CD4 T cells have been shown to be potent producers of gamma-interferon (IFN-γ), which, in turn, has been shown to trigger the anti-mycobacterial effects of macrophages in mice. While the role of IFN-γ in humans is less clear, studies have shown that 1,25-dihydroxy-vitamin D3, either alone or in combination with IFN-γ or tumor necrosis factor-alpha, activates human macrophages to inhibit *M. tuberculosis* infection. Furthermore, it is known that IFN-γ stimulates human macrophages to make 1,25-dihydroxy-vitamin D3. Similarly, IL-12 has been shown to play a role in stimulating resistance to *M. tuberculosis* infection. For a review of the immunology of *M. tuberculosis* infection see Chan and Kaufmann, in *Tuberculosis: Pathogenesis, Protection and Control*, Bloom (ed.), ASM Press, Washington, DC, 1994.

Accordingly, there is a need in the art for improved diagnostic methods for detecting tuberculosis. The present invention fulfills this need and further provides other related advantages.

SUMMARY OF THE INVENTION

Briefly stated, the present invention provides compositions and methods for diagnosing tuberculosis. In one aspect, polypeptides are provided comprising an antigenic portion of a soluble *M. tuberculosis* antigen, or a variant of such an antigen that differs only in conservative substitutions and/or modifications. In one embodiment of this aspect, the soluble antigen has one of the following N-terminal sequences:

(a) Asp-Pro-Val-Asp-Ala-Val-Ile-Asn-Thr-Thr-Cys-Asn-Tyr-Gly-Gln-Val-Val-Ala-Ala-Leu (SEQ ID NO: 115);

- Ala-Val-Glu-Ser-Gly-Met-Leu-Ala-Leu-Gly-Thr-Pro-Ala-Pro-Ser (b) (SEQ ID NO: 116); Ala-Ala-Met-Lys-Pro-Arg-Thr-Gly-Asp-Gly-Pro-Leu-Glu-Ala-Ala-(c) Lys-Glu-Gly-Arg (SEQ ID NO: 117); Tyr-Tyr-Trp-Cys-Pro-Gly-Gln-Pro-Phe-Asp-Pro-Ala-Trp-Gly-Pro(d) 5 (SEQ ID NO: 118); Asp-Ile-Gly-Ser-Glu-Ser-Thr-Glu-Asp-Gln-Gln-Xaa-Ala-Val (SEQ ID (e) NO: 119); Ala-Glu-Glu-Ser-Ile-Ser-Thr-Xaa-Glu-Xaa-Ile-Val-Pro (SEQ ID (f) NO: 120); 10 Asp-Pro-Glu-Pro-Ala-Pro-Pro-Val-Pro-Thr-Thr-Ala-Ala-Ser-Pro-Pro-(g) Ser (SEQ ID NO: 121); Ala-Pro-Lys-Thr-Tyr-Xaa-Glu-Glu-Leu-Lys-Gly-Thr-Asp-Thr-Gly (h) (SEQ ID NO: 122); Asp-Pro-Ala-Ser-Ala-Pro-Asp-Val-Pro-Thr-Ala-Ala-Gln-Leu-Thr-Ser-(i) 15 Leu-Leu-Asn-Ser-Leu-Ala-Asp-Pro-Asn-Val-Ser-Phe-Ala-Asn (SEQ ID NO: 123); Xaa-Asp-Ser-Glu-Lys-Ser-Ala-Thr-Ile-Lys-Val-Thr-Asp-Ala-Ser;(j) (SEQ ID NO: 129) Ala-Gly-Asp-Thr-Xaa-Ile-Tyr-Ile-Val-Gly-Asn-Leu-Thr-Ala-Asp; (k) 20 (SEQ ID NO: 130) or Ala-Pro-Glu-Ser-Gly-Ala-Gly-Leu-Gly-Gly-Thr-Val-Gln-Ala-Gly; **(l)** (SEQ ID NO: 131)
- wherein Xaa may be any amino acid.

In a related aspect, polypeptides are provided comprising an immunogenic portion of an *M. tuberculosis* antigen, or a variant of such an antigen that differs only in conservative substitutions and/or modifications, the antigen having one of the following N-terminal sequences:

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- (m) Xaa-Tyr-Ile-Ala-Tyr-Xaa-Thr-Thr-Ala-Gly-Ile-Val-Pro-Gly-Lys-Ile-Asn-Val-His-Leu-Val; (SEQ ID NO: 132) or
- (n) Asp-Pro-Pro-Asp-Pro-His-Gln-Xaa-Asp-Met-Thr-Lys-Gly-Tyr-Tyr-Pro-Gly-Gly-Arg-Arg-Xaa-Phe; (SEQ ID NO: 124)
- 5 wherein Xaa may be any amino acid.

In another embodiment, the soluble *M. tuberculosis* antigen comprises an amino acid sequence encoded by a DNA sequence selected from the group consisting of the sequences recited in SEQ ID NOS: 1, 2, 4-10, 13-25, 52, 94 and 96, the complements of said sequences, and DNA sequences that hybridize to a sequence recited in SEQ ID NOS: 1, 2, 4-10, 13-25, 52, 94 and 96 or a complement thereof under moderately stringent conditions.

In a related aspect, the polypeptides comprise an antigenic portion of a *M. tuberculosis* antigen, or a variant of such an antigen that differs only in conservative substitutions and/or modifications, wherein the antigen comprises an amino acid sequence encoded by a DNA sequence selected from the group consisting of the sequences recited in SEQ ID NOS: 26-51, 133, 134, 158-178 and 196, the complements of said sequences, and DNA sequences that hybridize to a sequence recited in SEQ ID NOS: 26-51, 133, 134, 158-178 and 196 or a complement thereof under moderately stringent conditions.

In related aspects, DNA sequences encoding the above polypeptides, recombinant expression vectors comprising these DNA sequences and host cells transformed or transfected with such expression vectors are also provided.

In another aspect, the present invention provides fusion proteins comprising a first and a second inventive polypeptide or, alternatively, an inventive polypeptide and a known *M. tuberculosis* antigen.

In further aspects of the subject invention, methods and diagnostic kits are provided for detecting tuberculosis in a patient. The methods comprise: (a) contacting a biological sample with at least one of the above polypeptides; and (b) detecting in the sample the presence of antibodies that bind to the polypeptide or polypeptides, thereby detecting *M. tuberculosis* infection in the biological sample. Suitable biological samples include whole blood, sputum, serum, plasma, saliva, cerebrospinal fluid and urine. The diagnostic kits comprise one or more of the above polypeptides in combination with a detection reagent.

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The present invention also provides methods for detecting *M. tuberculosis* infection comprising: (a) obtaining a biological sample from a patient; (b) contacting the sample with at least one oligonucleotide primer in a polymerase chain reaction, the oligonucleotide primer being specific for a DNA sequence encoding the above polypeptides; and (c) detecting in the sample a DNA sequence that amplifies in the presence of the first and second oligonucleotide primers. In one embodiment, the oligonucleotide primer comprises at least about 10 contiguous nucleotides of such a DNA sequence.

In a further aspect, the present invention provides a method for detecting *M. tuberculosis* infection in a patient comprising: (a) obtaining a biological sample from the patient; (b) contacting the sample with an oligonucleotide probe specific for a DNA sequence encoding the above polypeptides; and (c) detecting in the sample a DNA sequence that hybridizes to the oligonucleotide probe. In one embodiment, the oligonucleotide probe comprises at least about 15 contiguous nucleotides of such a DNA sequence.

In yet another aspect, the present invention provides antibodies, both polyclonal and monoclonal, that bind to the polypeptides described above, as well as methods for their use in the detection of *M. tuberculosis* infection.

These and other aspects of the present invention will become apparent upon reference to the following detailed description and attached drawings. All references disclosed herein are hereby incorporated by reference in their entirety as if each was incorporated individually.

BRIEF DESCRIPTION OF THE DRAWINGS AND SEQUENCE IDENTIFIERS

Figure 1A and B illustrate the stimulation of proliferation and interferon- γ production in T cells derived from a first and a second *M. tuberculosis*-immune donor, respectively, by the 14 Kd, 20 Kd and 26 Kd antigens described in Example 1.

Figures 2A-D illustrate the reactivity of antisera raised against secretory *M. tuberculosis* proteins, the known *M. tuberculosis* antigen 85b and the inventive antigens Tb38-1 and TbH-9, respectively, with *M. tuberculosis* lysate (lane 2), *M. tuberculosis* secretory proteins (lane 3), recombinant Tb38-1 (lane 4), recombinant TbH-9 (lane 5) and recombinant 85b (lane 5).

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Figure 3A illustrates the stimulation of proliferation in a TbH-9-specific T cell clone by secretory *M. tuberculosis* proteins, recombinant TbH-9 and a control antigen, TbRa11.

Figure 3B illustrates the stimulation of interferon-γ production in a TbH-9-specific T cell clone by secretory *M. tuberculosis* proteins, PPD and recombinant TbH-9.

Figure 4 illustrates the reactivity of two representative polypeptides with sera from *M. tuberculosis*-infected and uninfected individuals, as compared to the reactivity of bacterial lysate.

Figure 5 shows the reactivity of four representative polypeptides with sera from *M. tuberculosis*-infected and uninfected individuals, as compared to the reactivity of the 38 kD antigen.

Figure 6 shows the reactivity of recombinant 38 kD and TbRall antigens with sera from *M. tuberculosis* patients, PPD positive donors and normal donors.

Figure 7 shows the reactivity of the antigen TbRa2A with 38 kD negative sera.

Figure 8 shows the reactivity of the antigen of SEQ ID NO: 60 with sera from M. tuberculosis patients and normal donors.

Figure 9 illustrates the reactivity of the recombinant antigen TbH-29 (SEQ ID NO: 137) with sera from *M. tuberculosis* patients, PPD positive donors and normal donors as determined by indirect ELISA.

Figure 10 illustrates the reactivity of the recombinant antigen TbH-33 (SEQ ID NO: 140) with sera from *M. tuberculosis* patients and from normal donors, and with a pool of sera from *M. tuberculosis* patients, as determined both by direct and indirect ELISA

Figure 11 illustrates the reactivity of increasing concentrations of the recombinant antigen TbH-33 (SEQ ID NO: 140) with sera from M. tuberculosis patients and from normal donors as determined by ELISA.

SEQ. ID NO. 1 is the DNA sequence of TbRa1.

SEQ. ID NO. 2 is the DNA sequence of TbRa10.

SEQ. ID NO. 3 is the DNA sequence of TbRall.

30 SEQ. ID NO. 4 is the DNA sequence of TbRa12.

	SEQ. ID NO. 5 is the DNA sequence of TbRa13.
	SEQ. ID NO. 6 is the DNA sequence of TbRa16.
	SEQ. ID NO. 7 is the DNA sequence of TbRa17.
	SEQ. ID NO. 8 is the DNA sequence of TbRa18.
5	SEQ. ID NO. 9 is the DNA sequence of TbRa19.
	SEQ. ID NO. 10 is the DNA sequence of TbRa24.
	SEQ. ID NO. 11 is the DNA sequence of TbRa26.
	SEQ. ID NO. 12 is the DNA sequence of TbRa28.
	SEQ. ID NO. 13 is the DNA sequence of TbRa29.
10	SEQ. ID NO. 14 is the DNA sequence of TbRa2A.
	SEQ. ID NO. 15 is the DNA sequence of TbRa3.
	SEQ. ID NO. 16 is the DNA sequence of TbRa32.
	SEQ. ID NO. 17 is the DNA sequence of TbRa35.
	SEQ. ID NO. 18 is the DNA sequence of TbRa36.
15	SEQ. ID NO. 19 is the DNA sequence of TbRa4.
	SEQ. ID NO. 20 is the DNA sequence of TbRa9.
	SEQ. ID NO. 21 is the DNA sequence of TbRaB.
	SEQ. ID NO. 22 is the DNA sequence of TbRaC.
	SEQ. ID NO. 23 is the DNA sequence of TbRaD.
20	SEQ. ID NO. 24 is the DNA sequence of YYWCPG.
	SEQ. ID NO. 25 is the DNA sequence of AAMK.
	SEQ. ID NO. 26 is the DNA sequence of TbL-23.
	SEQ. ID NO. 27 is the DNA sequence of TbL-24.
	SEQ. ID NO. 28 is the DNA sequence of TbL-25.
25	SEQ. ID NO. 29 is the DNA sequence of TbL-28.
	SEQ. ID NO. 30 is the DNA sequence of TbL-29.
	SEQ. ID NO. 31 is the DNA sequence of TbH-5.
	SEQ. ID NO. 32 is the DNA sequence of TbH-8.
	SEQ. ID NO. 33 is the DNA sequence of TbH-9.
30	SEQ. ID NO. 34 is the DNA sequence of TbM-1.

	SEQ. ID NO. 35 is the DNA sequence of 1 bM-3.
	SEQ. ID NO. 36 is the DNA sequence of TbM-6.
	SEQ. ID NO. 37 is the DNA sequence of TbM-7.
	SEQ. ID NO. 38 is the DNA sequence of TbM-9.
5	SEQ. ID NO. 39 is the DNA sequence of TbM-12.
	SEQ. ID NO. 40 is the DNA sequence of TbM-13.
	SEQ. ID NO. 41 is the DNA sequence of TbM-14.
	SEQ. ID NO. 42 is the DNA sequence of TbM-15.
	SEQ. ID NO. 43 is the DNA sequence of TbH-4.
10	SEQ. ID NO. 44 is the DNA sequence of TbH-4-FWD.
	SEQ. ID NO. 45 is the DNA sequence of TbH-12.
	SEQ. ID NO. 46 is the DNA sequence of Tb38-1.
	SEQ. ID NO. 47 is the DNA sequence of Tb38-4.
	SEQ. ID NO. 48 is the DNA sequence of TbL-17.
15	SEQ. ID NO. 49 is the DNA sequence of TbL-20.
	SEQ. ID NO. 50 is the DNA sequence of TbL-21.
	SEQ. ID NO. 51 is the DNA sequence of TbH-16.
	SEQ. ID NO. 52 is the DNA sequence of DPEP.
	SEQ. ID NO. 53 is the deduced amino acid sequence of DPEP.
20	SEQ. ID NO. 54 is the protein sequence of DPV N-terminal Antigen.
	SEQ. ID NO. 55 is the protein sequence of AVGS N-terminal Antigen.
	SEQ. ID NO. 56 is the protein sequence of AAMK N-terminal Antigen.
	SEQ. ID NO. 57 is the protein sequence of YYWC N-terminal Antigen.
	SEQ. ID NO. 58 is the protein sequence of DIGS N-terminal Antigen.
25	SEQ. ID NO. 59 is the protein sequence of AEES N-terminal Antigen.
	SEQ. ID NO. 60 is the protein sequence of DPEP N-terminal Antigen.
	SEQ. ID NO. 61 is the protein sequence of APKT N-terminal Antigen.
	SEQ. ID NO. 62 is the protein sequence of DPAS N-terminal Antigen.
	SEQ. ID NO. 63 is the deduced amino acid sequence of TbM-1 Peptide.
30	SEO ID NO. 64 is the deduced amino acid sequence of TbRa1.

	SEQ. 1D NO. 65 is the deduced amino acid sequence of Tokato.
	SEQ. ID NO. 66 is the deduced amino acid sequence of TbRa11.
	SEQ. ID NO. 67 is the deduced amino acid sequence of TbRa12.
	SEQ. ID NO. 68 is the deduced amino acid sequence of TbRa13.
5	SEQ. ID NO. 69 is the deduced amino acid sequence of TbRa16.
	SEQ. ID NO. 70 is the deduced amino acid sequence of TbRa17.
	SEQ. ID NO. 71 is the deduced amino acid sequence of TbRa18.
	SEQ. ID NO. 72 is the deduced amino acid sequence of TbRa19.
	SEQ. ID NO. 73 is the deduced amino acid sequence of TbRa24.
10	SEQ. ID NO. 74 is the deduced amino acid sequence of TbRa26.
	SEQ. ID NO. 75 is the deduced amino acid sequence of TbRa28.
	SEQ. ID NO. 76 is the deduced amino acid sequence of TbRa29.
	SEQ. ID NO. 77 is the deduced amino acid sequence of TbRa2A.
	SEQ. ID NO. 78 is the deduced amino acid sequence of TbRa3.
15	SEQ. ID NO. 79 is the deduced amino acid sequence of TbRa32.
	SEQ. ID NO. 80 is the deduced amino acid sequence of TbRa35.
	SEQ. ID NO. 81 is the deduced amino acid sequence of TbRa36.
	SEQ. ID NO. 82 is the deduced amino acid sequence of TbRa4.
	SEQ. ID NO. 83 is the deduced amino acid sequence of TbRa9.
20	SEQ. ID NO. 84 is the deduced amino acid sequence of TbRaB.
	SEQ. ID NO. 85 is the deduced amino acid sequence of TbRaC.
	SEQ. ID NO. 86 is the deduced amino acid sequence of TbRaD.
	SEQ. ID NO. 87 is the deduced amino acid sequence of YYWCPG.
	SEQ. ID NO. 88 is the deduced amino acid sequence of TbAAMK.
25	SEQ. ID NO. 89 is the deduced amino acid sequence of Tb38-1.
	SEQ. ID NO. 90 is the deduced amino acid sequence of TbH-4.
	SEQ. ID NO. 91 is the deduced amino acid sequence of TbH-8.
	SEQ. ID NO. 92 is the deduced amino acid sequence of TbH-9.
	SEQ. ID NO. 93 is the deduced amino acid sequence of TbH-12.
30	SEQ. ID NO. 94 is the DNA sequence of DPAS.

SEO. ID NO. 95 is the deduced amino acid sequence of DPAS. SEQ. ID NO. 96 is the DNA sequence of DPV. SEQ. ID NO. 97 is the deduced amino acid sequence of DPV. SEQ. ID NO. 98 is the DNA sequence of ESAT-6. SEQ. ID NO. 99 is the deduced amino acid sequence of ESAT-6. 5 SEQ. ID NO. 100 is the DNA sequence of TbH-8-2. SEO. ID NO. 101 is the DNA sequence of TbH-9FL. SEQ. ID NO. 102 is the deduced amino acid sequence of TbH-9FL. SEO. ID NO. 103 is the DNA sequence of TbH-9-1. SEQ. ID NO. 104 is the deduced amino acid sequence of TbH-9-1. 10 SEO. ID NO. 105 is the DNA sequence of TbH-9-4. SEO. ID NO. 106 is the deduced amino acid sequence of TbH-9-4. SEQ. ID NO. 107 is the DNA sequence of Tb38-1F2 IN. SEO. ID NO. 108 is the DNA sequence of Tb38-1F2 RP. SEO. ID NO. 109 is the deduced amino acid sequence of Tb37-FL. 15 SEQ. ID NO. 110 is the deduced amino acid sequence of Tb38-IN. SEO. ID NO. 111 is the DNA sequence of Tb38-1F3. SEQ. ID NO. 112 is the deduced amino acid sequence of Tb38-1F3. SEO. ID NO. 113 is the DNA sequence of Tb38-1F5. SEO, ID NO. 114 is the DNA sequence of Tb38-1F6. 20 SEQ. ID NO. 115 is the deduced N-terminal amino acid sequence of DPV. SEQ. ID NO. 116 is the deduced N-terminal amino acid sequence of AVGS. SEQ. ID NO. 117 is the deduced N-terminal amino acid sequence of AAMK. SEQ. ID NO. 118 is the deduced N-terminal amino acid sequence of YYWC. SEQ. ID NO. 119 is the deduced N-terminal amino acid sequence of DIGS. 25 SEQ. ID NO. 120 is the deduced N-terminal amino acid sequence of AAES. SEQ. ID NO. 121 is the deduced N-terminal amino acid sequence of DPEP. SEQ. ID NO. 122 is the deduced N-terminal amino acid sequence of APKT. SEQ. ID NO. 123 is the deduced N-terminal amino acid sequence of DPAS.

SEQ. ID NO. 124 is the protein sequence of DPPD N-terminal Antigen.

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SEQ ID NO. 125-128 are the protein sequences of four DPPD cyanogen bromide fragments.

SEQ ID NO. 129 is the N-terminal protein sequence of XDS antigen.

SEQ ID NO. 130 is the N-terminal protein sequence of AGD antigen.

5 SEQ ID NO. 131 is the N-terminal protein sequence of APE antigen.

SEQ ID NO. 132 is the N-terminal protein sequence of XYI antigen.

SEQ ID NO. 133 is the DNA sequence of TbH-29.

SEQ ID NO. 134 is the DNA sequence of TbH-30.

SEQ ID NO. 135 is the DNA sequence of TbH-32.

SEQ ID NO. 136 is the DNA sequence of TbH-33.

SEQ ID NO. 137 is the predicted amino acid sequence of TbH-29.

SEQ ID NO. 138 is the predicted amino acid sequence of TbH-30.

SEQ ID NO. 139 is the predicted amino acid sequence of TbH-32.

SEQ ID NO. 140 is the predicted amino acid sequence of TbH-33.

SEQ ID NO: 141-146 are PCR primers used in the preparation of a fusion protein containing TbRa3, 38 kD and Tb38-1.

SEQ ID NO: 147 is the DNA sequence of the fusion protein containing TbRa3, 38 kD and Tb38-1.

SEQ ID NO: 148 is the amino acid sequence of the fusion protein containing TbRa3,

20 38 kD and Tb38-1.

SEQ ID NO: 149 is the DNA sequence of the M. tuberculosis antigen 38 kD.

SEQ ID NO: 150 is the amino acid sequence of the M. tuberculosis antigen 38 kD.

SEQ ID NO: 151 is the DNA sequence of XP14.

SEQ ID NO: 152 is the DNA sequence of XP24.

25 SEQ ID NO: 153 is the DNA sequence of XP31.

SEQ ID NO: 154 is the 5' DNA sequence of XP32.

SEQ ID NO: 155 is the 3' DNA sequence of XP32.

SEQ ID NO: 156 is the predicted amino acid sequence of XP14.

SEQ ID NO: 157 is the predicted amino acid sequence encoded by the reverse

30 complement of XP14.

	SEQ ID NO: 158 is the DNA sequence of XP27.
	SEQ ID NO: 159 is the DNA sequence of XP36.
	SEQ ID NO: 160 is the 5' DNA sequence of XP4.
	SEQ ID NO: 161 is the 5' DNA sequence of XP5.
5	SEQ ID NO: 162 is the 5' DNA sequence of XP17.
	SEQ ID NO: 163 is the 5' DNA sequence of XP30.
	SEQ ID NO: 164 is the 5' DNA sequence of XP2.
	SEQ ID NO: 165 is the 3' DNA sequence of XP2.
	SEQ ID NO: 166 is the 5' DNA sequence of XP3.
10	SEQ ID NO: 167 is the 3' DNA sequence of XP3.
	SEQ ID NO: 168 is the 5' DNA sequence of XP6.
	SEQ ID NO: 169 is the 3' DNA sequence of XP6.
	SEQ ID NO: 170 is the 5' DNA sequence of XP18.
	SEQ ID NO: 171 is the 3' DNA sequence of XP18.
15	SEQ ID NO: 172 is the 5' DNA sequence of XP19.
	SEQ ID NO: 173 is the 3' DNA sequence of XP19.
	SEQ ID NO: 174 is the 5' DNA sequence of XP22.
	SEQ ID NO: 175 is the 3' DNA sequence of XP22.
	SEQ ID NO: 176 is the 5' DNA sequence of XP25.
20	SEQ ID NO: 177 is the 3' DNA sequence of XP25.
	SEQ ID NO: 178 is the full-length DNA sequence of TbH4-XP1.
	SEQ ID NO: 179 is the predicted amino acid sequence of TbH4-XP1.
	SEQ ID NO: 180 is the predicted amino acid sequence encoded by the reverse
	complement of TbH4-XP1.
25	SEQ ID NO: 181 is a first predicted amino acid sequence encoded by XP36.
	SEQ ID NO: 182 is a second predicted amino acid sequence encoded by XP36.
	SEQ ID NO: 183 is the predicted amino acid sequence encoded by the reverse
	complement of XP36.
	SEQ ID NO: 184 is the DNA sequence of RDIF2.
30	SEQ ID NO: 185 is the DNA sequence of RDIF5.

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SEQ ID NO: 186 is the DNA sequence of RDIF8.

SEQ ID NO: 187 is the DNA sequence of RDIF10.

SEQ ID NO: 188 is the DNA sequence of RDIF11.

SEQ ID NO: 189 is the predicted amino acid sequence of RDIF2.

SEQ ID NO: 190 is the predicted amino acid sequence of RDIF5.

SEQ ID NO: 191 is the predicted amino acid sequence of RDIF8.

SEQ ID NO: 192 is the predicted amino acid sequence of RDIF10.

SEQ ID NO: 193 is the predicted amino acid sequence of RDIF11.

SEQ ID NO: 194 is the 5' DNA sequence of RDIF12.

SEQ ID NO: 195 is the 3' DNA sequence of RDIF12.

SEQ ID NO: 196 is the DNA sequence of RDIF7.

SEQ ID NO: 197 is the predicted amino acid sequence of RDIF7.

SEQ ID NO: 198 is the DNA sequence of DIF2-1.

SEQ ID NO: 199 is the predicted amino acid sequence of DIF2-1.

SEQ ID NO: 200-207 are PCR primers used in the preparation of a fusion protein containing TbRa3, 38 kD, Tb38-1 and DPEP (hereinafter referred to as TbF-2).

SEQ ID NO: 208 is the DNA sequence of the fusion protein TbF-2.

SEQ ID NO: 209 is the amino acid sequence of the fusion protein TbF-2.

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DETAILED DESCRIPTION OF THE INVENTION

As noted above, the present invention is generally directed to compositions and methods for diagnosing tuberculosis. The compositions of the subject invention include polypeptides that comprise at least one antigenic portion of a *M. tuberculosis* antigen, or a variant of such an antigen that differs only in conservative substitutions and/or modifications. Polypeptides within the scope of the present invention include, but are not limited to, soluble *M. tuberculosis* antigens. A "soluble *M. tuberculosis* antigen" is a protein of *M. tuberculosis* origin that is present in *M. tuberculosis* culture filtrate. As used herein, the term "polypeptide" encompasses amino acid chains of any length, including full length proteins (i.e., antigens), wherein the amino acid residues are linked by covalent peptide bonds. Thus,

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a polypeptide comprising an antigenic portion of one of the above antigens may consist entirely of the antigenic portion, or may contain additional sequences. The additional sequences may be derived from the native *M. tuberculosis* antigen or may be heterologous, and such sequences may (but need not) be antigenic.

An "antigenic portion" of an antigen (which may or may not be soluble) is a portion that is capable of reacting with sera obtained from an *M. tuberculosis*-infected individual (*i.e.*, generates an absorbance reading with sera from infected individuals that is at least three standard deviations above the absorbance obtained with sera from uninfected individuals, in a representative ELISA assay described herein). An "*M. tuberculosis*-infected individual" is a human who has been infected with *M. tuberculosis* (*e.g.*, has an intradermal skin test response to PPD that is at least 0.5 cm in diameter). Infected individuals may display symptoms of tuberculosis or may be free of disease symptoms. Polypeptides comprising at least an antigenic portion of one or more *M. tuberculosis* antigens as described herein may generally be used, alone or in combination, to detect tuberculosis in a patient.

The compositions and methods of this invention also encompass variants of the above polypeptides. A "variant," as used herein, is a polypeptide that differs from the native antigen only in conservative substitutions and/or modifications, such that the antigenic properties of the polypeptide are retained. Such variants may generally be identified by modifying one of the above polypeptide sequences, and evaluating the antigenic properties of the modified polypeptide using, for example, the representative procedures described herein.

A "conservative substitution" is one in which an amino acid is substituted for another amino acid that has similar properties, such that one skilled in the art of peptide chemistry would expect the secondary structure and hydropathic nature of the polypeptide to be substantially unchanged. In general, the following groups of amino acids represent conservative changes: (1) ala, pro, gly, glu, asp, gln, asn, ser, thr; (2) cys, ser, tyr, thr; (3) val, ile, leu, met, ala, phe; (4) lys, arg, his; and (5) phe, tyr, trp, his.

Variants may also (or alternatively) be modified by, for example, the deletion or addition of amino acids that have minimal influence on the antigenic properties, secondary structure and hydropathic nature of the polypeptide. For example, a polypeptide may be conjugated to a signal (or leader) sequence at the N-terminal end of the protein which co-

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translationally or post-translationally directs transfer of the protein. The polypeptide may also be conjugated to a linker or other sequence for ease of synthesis, purification or identification of the polypeptide (e.g., poly-His), or to enhance binding of the polypeptide to a solid support. For example, a polypeptide may be conjugated to an immunoglobulin Fc region.

In a related aspect, combination polypeptides are disclosed. A "combination polypeptide" is a polypeptide comprising at least one of the above antigenic portions and one or more additional antigenic *M. tuberculosis* sequences, which are joined via a peptide linkage into a single amino acid chain. The sequences may be joined directly (*i.e.*, with no intervening amino acids) or may be joined by way of a linker sequence (*e.g.*, Gly-Cys-Gly) that does not significantly diminish the antigenic properties of the component polypeptides.

In general, *M. tuberculosis* antigens, and DNA sequences encoding such antigens, may be prepared using any of a variety of procedures. For example, soluble antigens may be isolated from *M. tuberculosis* culture filtrate by procedures known to those of ordinary skill in the art, including anion-exchange and reverse phase chromatography. Purified antigens may then be evaluated for a desired property, such as the ability to react with sera obtained from an *M. tuberculosis*-infected individual. Such screens may be performed using the representative methods described herein. Antigens may then be partially sequenced using, for example, traditional Edman chemistry. *See* Edman and Berg, *Eur. J. Biochem.* 80:116-132, 1967.

Antigens may also be produced recombinantly using a DNA sequence that encodes the antigen, which has been inserted into an expression vector and expressed in an appropriate host. DNA molecules encoding soluble antigens may be isolated by screening an appropriate *M. tuberculosis* expression library with anti-sera (e.g., rabbit) raised specifically against soluble *M. tuberculosis* antigens. DNA sequences encoding antigens that may or may not be soluble may be identified by screening an appropriate *M. tuberculosis* genomic or cDNA expression library with sera obtained from patients infected with *M. tuberculosis*. Such screens may generally be performed using techniques well known in the art, such as those described in Sambrook et al., *Molecular Cloning: A Laboratory Manual*, Cold Spring Harbor Laboratories, Cold Spring Harbor, NY, 1989.

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DNA sequences encoding soluble antigens may also be obtained by screening an appropriate *M. tuberculosis* cDNA or genomic DNA library for DNA sequences that hybridize to degenerate oligonucleotides derived from partial amino acid sequences of isolated soluble antigens. Degenerate oligonucleotide sequences for use in such a screen may be designed and synthesized, and the screen may be performed, as described (for example) in Sambrook et al., *Molecular Cloning: A Laboratory Manual*, Cold Spring Harbor Laboratories, Cold Spring Harbor, NY (and references cited therein). Polymerase chain reaction (PCR) may also be employed, using the above oligonucleotides in methods well known in the art, to isolate a nucleic acid probe from a cDNA or genomic library. The library screen may then be performed using the isolated probe.

Regardless of the method of preparation, the antigens described herein are "antigenic." More specifically, the antigens have the ability to react with sera obtained from an *M. tuberculosis*-infected individual. Reactivity may be evaluated using, for example, the representative ELISA assays described herein, where an absorbance reading with sera from infected individuals that is at least three standard deviations above the absorbance obtained with sera from uninfected individuals is considered positive.

Antigenic portions of *M. tuberculosis* antigens may be prepared and identified using well known techniques, such as those summarized in Paul, *Fundamental Immunology*, 3d ed., Raven Press, 1993, pp. 243-247 and references cited therein. Such techniques include screening polypeptide portions of the native antigen for antigenic properties. The representative ELISAs described herein may generally be employed in these screens. An antigenic portion of a polypeptide is a portion that, within such representative assays, generates a signal in such assays that is substantially similar to that generated by the full length antigen. In other words, an antigenic portion of a *M. tuberculosis* antigen generates at least about 20%, and preferably about 100%, of the signal induced by the full length antigen in a model ELISA as described herein.

Portions and other variants of *M. tuberculosis* antigens may be generated by synthetic or recombinant means. Synthetic polypeptides having fewer than about 100 amino acids, and generally fewer than about 50 amino acids, may be generated using techniques well known in the art. For example, such polypeptides may be synthesized using any of the

commercially available solid-phase techniques, such as the Merrifield solid-phase synthesis method, where amino acids are sequentially added to a growing amino acid chain. See Merrifield, J. Am. Chem. Soc. 85:2149-2146, 1963. Equipment for automated synthesis of polypeptides is commercially available from suppliers such as Applied BioSystems, Inc., Foster City, CA, and may be operated according to the manufacturer's instructions. Variants of a native antigen may generally be prepared using standard mutagenesis techniques, such as oligonucleotide-directed site-specific mutagenesis. Sections of the DNA sequence may also be removed using standard techniques to permit preparation of truncated polypeptides.

Recombinant polypeptides containing portions and/or variants of a native antigen may be readily prepared from a DNA sequence encoding the polypeptide using a variety of techniques well known to those of ordinary skill in the art. For example, supernatants from suitable host/vector systems which secrete recombinant protein into culture media may be first concentrated using a commercially available filter. Following concentration, the concentrate may be applied to a suitable purification matrix such as an affinity matrix or an ion exchange resin. Finally, one or more reverse phase HPLC steps can be employed to further purify a recombinant protein.

Any of a variety of expression vectors known to those of ordinary skill in the art may be employed to express recombinant polypeptides as described herein. Expression may be achieved in any appropriate host cell that has been transformed or transfected with an expression vector containing a DNA molecule that encodes a recombinant polypeptide. Suitable host cells include prokaryotes, yeast and higher eukaryotic cells. Preferably, the host cells employed are *E. coli*, yeast or a mammalian cell line, such as COS or CHO. The DNA sequences expressed in this manner may encode naturally occurring antigens, portions of naturally occurring antigens, or other variants thereof.

In general, regardless of the method of preparation, the polypeptides disclosed herein are prepared in substantially pure form. Preferably, the polypeptides are at least about 80% pure, more preferably at least about 90% pure and most preferably at least about 99% pure. For use in the methods described herein, however, such substantially pure polypeptides may be combined.

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In certain specific embodiments, the subject invention discloses polypeptides comprising at least an antigenic portion of a soluble *M. tuberculosis* antigen (or a variant of such an antigen), where the antigen has one of the following N-terminal sequences:

- (a) Asp-Pro-Val-Asp-Ala-Val-Ile-Asn-Thr-Thr-Cys-Asn-Tyr-Gly-Gln-Val-Val-Ala-Ala-Leu (SEQ ID NO: 115);
- (b) Ala-Val-Glu-Ser-Gly-Met-Leu-Ala-Leu-Gly-Thr-Pro-Ala-Pro-Ser (SEQ ID NO: 116);
- (c) Ala-Ala-Met-Lys-Pro-Arg-Thr-Gly-Asp-Gly-Pro-Leu-Glu-Ala-Ala-Lys-Glu-Gly-Arg (SEQ ID NO: 117);
- (d) Tyr-Tyr-Trp-Cys-Pro-Gly-Gln-Pro-Phe-Asp-Pro-Ala-Trp-Gly-Pro (SEQ ID NO: 118);
 - (e) Asp-Ile-Gly-Ser-Glu-Ser-Thr-Glu-Asp-Gln-Gln-Xaa-Ala-Val (SEQ ID NO: 119);
 - (f) Ala-Glu-Glu-Ser-Ile-Ser-Thr-Xaa-Glu-Xaa-Ile-Val-Pro (SEQ ID NO: 120);
 - (g) Asp-Pro-Glu-Pro-Ala-Pro-Pro-Val-Pro-Thr-Thr-Ala-Ala-Ser-Pro-Pro-Ser (SEQ ID NO: 121);
 - (h) Ala-Pro-Lys-Thr-Tyr-Xaa-Glu-Glu-Leu-Lys-Gly-Thr-Asp-Thr-Gly (SEQ ID NO: 122);
 - (i) Asp-Pro-Ala-Ser-Ala-Pro-Asp-Val-Pro-Thr-Ala-Ala-Gln-Gln-Thr-Ser-Leu-Leu-Asn-Ser-Leu-Ala-Asp-Pro-Asn-Val-Ser-Phe-Ala-Asn (SEQ ID NO: 123);
 - (j) Xaa-Asp-Ser-Glu-Lys-Ser-Ala-Thr-Ile-Lys-Val-Thr-Asp-Ala-Ser; (SEQ ID NO: 129)
 - (k) Ala-Gly-Asp-Thr-Xaa-Ile-Tyr-Ile-Val-Gly-Asn-Leu-Thr-Ala-Asp; (SEQ ID NO: 130) or
 - (l) Ala-Pro-Glu-Ser-Gly-Ala-Gly-Leu-Gly-Gly-Thr-Val-Gln-Ala-Gly; (SEQ ID NO: 131)

wherein Xaa may be any amino acid, preferably a cysteine residue. A DNA sequence an encoding the antigen identified as (g) above is provided in SEQ ID NO: 52, the deduced

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amino acid sequence of which is provided in SEQ ID NO: 53. A DNA sequence encoding the antigen identified as (a) above is provided in SEQ ID NO: 96; its deduced amino acid sequence is provided in SEQ ID NO: 97. A DNA sequence corresponding to antigen (d) above is provided in SEQ ID NO: 24, a DNA sequence corresponding to antigen (c) is provided in SEQ ID NO: 25 and a DNA sequence corresponding to antigen (I) is disclosed in SEQ ID NO: 94 and its deduced amino acid sequence is provided in SEQ ID NO: 95.

In a further specific embodiment, the subject invention discloses polypeptides comprising at least an immunogenic portion of an *M. tuberculosis* antigen having one of the following N-terminal sequences, or a variant thereof that differs only in conservative substitutions and/or modifications:

- (m) Xaa-Tyr-Ile-Ala-Tyr-Xaa-Thr-Thr-Ala-Gly-Ile-Val-Pro-Gly-Lys-Ile-Asn-Val-His-Leu-Val; (SEQ ID NO: 132) or
- (n) Asp-Pro-Pro-Asp-Pro-His-Gln-Xaa-Asp-Met-Thr-Lys-Gly-Tyr-Tyr-Pro-Gly-Gly-Arg-Arg-Xaa-Phe; (SEQ ID NO: 124)

wherein Xaa may be any amino acid, preferably a cysteine residue.

In other specific embodiments, the subject invention discloses polypeptides comprising at least an antigenic portion of a soluble *M. tuberculosis* antigen (or a variant of such an antigen) that comprises one or more of the amino acid sequences encoded by (a) the DNA sequences of SEQ ID NOS: 1, 2, 4-10, 13-25, 52, 94 and 96, (b) the complements of such DNA sequences, or (c) DNA sequences substantially homologous to a sequence in (a) or (b).

In further specific embodiments, the subject invention discloses polypeptides comprising at least an antigenic portion of a *M. tuberculosis* antigen (or a variant of such an antigen), which may or may not be soluble, that comprises one or more of the amino acid sequences encoded by (a) the DNA sequences of SEQ ID NOS: 26-51, 133, 134, 158-178 and 196, (b) the complements of such DNA sequences or (c) DNA sequences substantially homologous to a sequence in (a) or (b).

In the specific embodiments discussed above, the *M. tuberculosis* antigens include variants that are encoded DNA sequences which are substantially homologous to one

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or more of DNA sequences specifically recited herein. "Substantial homology," as used herein, refers to DNA sequences that are capable of hybridizing under moderately stringent conditions. Suitable moderately stringent conditions include prewashing in a solution of 5X SSC, 0.5% SDS, 1.0 mM EDTA (pH 8.0); hybridizing at 50°C-65°C, 5X SSC, overnight or, in the event of cross-species homology, at 45°C with 0.5X SSC; followed by washing twice at 65°C for 20 minutes with each of 2X, 0.5X and 0.2X SSC containing 0.1% SDS). Such hybridizing DNA sequences are also within the scope of this invention, as are nucleotide sequences that, due to code degeneracy, encode an immunogenic polypeptide that is encoded by a hybridizing DNA sequence.

In a related aspect, the present invention provides fusion proteins comprising a first and a second inventive polypeptide or, alternatively, a polypeptide of the present invention and a known *M. tuberculosis* antigen, such as the 38 kD antigen described above or ESAT-6 (SEQ ID NOS: 98 and 99), together with variants of such fusion proteins. The fusion proteins of the present invention may also include a linker peptide between the first and second polypeptides.

A DNA sequence encoding a fusion protein of the present invention is constructed using known recombinant DNA techniques to assemble separate DNA sequences encoding the first and second polypeptides into an appropriate expression vector. The 3' end of a DNA sequence encoding the first polypeptide is ligated, with or without a peptide linker, to the 5' end of a DNA sequence encoding the second polypeptide so that the reading frames of the sequences are in phase to permit mRNA translation of the two DNA sequences into a single fusion protein that retains the biological activity of both the first and the second polypeptides.

A peptide linker sequence may be employed to separate the first and the second polypeptides by a distance sufficient to ensure that each polypeptide folds into its secondary and tertiary structures. Such a peptide linker sequence is incorporated into the fusion protein using standard techniques well known in the art. Suitable peptide linker sequences may be chosen based on the following factors: (1) their ability to adopt a flexible extended conformation; (2) their inability to adopt a secondary structure that could interact with functional epitopes on the first and second polypeptides; and (3) the lack of hydrophobic

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or charged residues that might react with the polypeptide functional epitopes. Preferred peptide linker sequences contain Gly, Asn and Ser residues. Other near neutral amino acids, such as Thr and Ala may also be used in the linker sequence. Amino acid sequences which may be usefully employed as linkers include those disclosed in Maratea et al., *Gene 40*:39-46, 1985; Murphy et al., *Proc. Natl. Acad. Sci. USA 83*:8258-8562, 1986; U.S. Patent No. 4,935,233 and U.S. Patent No. 4,751,180. The linker sequence may be from 1 to about 50 amino acids in length. Peptide linker sequences are not required when the first and second polypeptides have non-essential N-terminal amino acid regions that can be used to separate the functional domains and prevent steric hindrance.

In another aspect, the present invention provides methods for using the polypeptides described above to diagnose tuberculosis. In this aspect, methods are provided for detecting *M. tuberculosis* infection in a biological sample, using one or more of the above polypeptides, alone or in combination. In embodiments in which multiple polypeptides are employed, polypeptides other than those specifically described herein, such as the 38 kD antigen described in Andersen and Hansen, *Infect. Immun.* 57:2481-2488, 1989, may be included. As used herein, a "biological sample" is any antibody-containing sample obtained from a patient. Preferably, the sample is whole blood, sputum, serum, plasma, saliva, cerebrospinal fluid or urine. More preferably, the sample is a blood, serum or plasma sample obtained from a patient or a blood supply. The polypeptide(s) are used in an assay, as described below, to determine the presence or absence of antibodies to the polypeptide(s) in the sample, relative to a predetermined cut-off value. The presence of such antibodies indicates previous sensitization to mycobacterial antigens which may be indicative of tuberculosis.

In embodiments in which more than one polypeptide is employed, the polypeptides used are preferably complementary (i.e., one component polypeptide will tend to detect infection in samples where the infection would not be detected by another component polypeptide). Complementary polypeptides may generally be identified by using each polypeptide individually to evaluate serum samples obtained from a series of patients known to be infected with *M. tuberculosis*. After determining which samples test positive (as described below) with each polypeptide, combinations of two or more polypeptides may be

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formulated that are capable of detecting infection in most, or all, of the samples tested. Such polypeptides are complementary. For example, approximately 25-30% of sera from tuberculosis-infected individuals are negative for antibodies to any single protein, such as the 38 kD antigen mentioned above. Complementary polypeptides may, therefore, be used in combination with the 38 kD antigen to improve sensitivity of a diagnostic test.

There are a variety of assay formats known to those of ordinary skill in the art for using one or more polypeptides to detect antibodies in a sample. See, e.g., Harlow and Lane, Antibodies: A Laboratory Manual, Cold Spring Harbor Laboratory, 1988, which is incorporated herein by reference. In a preferred embodiment, the assay involves the use of polypeptide immobilized on a solid support to bind to and remove the antibody from the sample. The bound antibody may then be detected using a detection reagent that contains a reporter group. Suitable detection reagents include antibodies that bind to the antibody/polypeptide complex and free polypeptide labeled with a reporter group (e.g., in a semi-competitive assay). Alternatively, a competitive assay may be utilized, in which an antibody that binds to the polypeptide is labeled with a reporter group and allowed to bind to the immobilized antigen after incubation of the antigen with the sample. The extent to which components of the sample inhibit the binding of the labeled antibody to the polypeptide is indicative of the reactivity of the sample with the immobilized polypeptide.

The solid support may be any solid material known to those of ordinary skill in the art to which the antigen may be attached. For example, the solid support may be a test well in a microtiter plate or a nitrocellulose or other suitable membrane. Alternatively, the support may be a bead or disc, such as glass, fiberglass, latex or a plastic material such as polystyrene or polyvinylchloride. The support may also be a magnetic particle or a fiber optic sensor, such as those disclosed, for example, in U.S. Patent No. 5,359,681.

The polypeptides may be bound to the solid support using a variety of techniques known to those of ordinary skill in the art, which are amply described in the patent and scientific literature. In the context of the present invention, the term "bound" refers to both noncovalent association, such as adsorption, and covalent attachment (which may be a direct linkage between the antigen and functional groups on the support or may be a linkage by way of a cross-linking agent). Binding by adsorption to a well in a microtiter plate or to a

membrane is preferred. In such cases, adsorption may be achieved by contacting the polypeptide, in a suitable buffer, with the solid support for a suitable amount of time. The contact time varies with temperature, but is typically between about 1 hour and 1 day. In general, contacting a well of a plastic microtiter plate (such as polystyrene or polyvinylchloride) with an amount of polypeptide ranging from about 10 ng to about 1 µg, and preferably about 100 ng, is sufficient to bind an adequate amount of antigen.

Covalent attachment of polypeptide to a solid support may generally be achieved by first reacting the support with a bifunctional reagent that will react with both the support and a functional group, such as a hydroxyl or amino group, on the polypeptide. For example, the polypeptide may be bound to supports having an appropriate polymer coating using benzoquinone or by condensation of an aldehyde group on the support with an amine and an active hydrogen on the polypeptide (see, e.g., Pierce Immunotechnology Catalog and Handbook, 1991, at A12-A13).

In certain embodiments, the assay is an enzyme linked immunosorbent assay (ELISA). This assay may be performed by first contacting a polypeptide antigen that has been immobilized on a solid support, commonly the well of a microtiter plate, with the sample, such that antibodies to the polypeptide within the sample are allowed to bind to the immobilized polypeptide. Unbound sample is then removed from the immobilized polypeptide and a detection reagent capable of binding to the immobilized antibody-polypeptide complex is added. The amount of detection reagent that remains bound to the solid support is then determined using a method appropriate for the specific detection reagent.

More specifically, once the polypeptide is immobilized on the support as described above, the remaining protein binding sites on the support are typically blocked. Any suitable blocking agent known to those of ordinary skill in the art, such as bovine serum albumin or Tween 20^{TM} (Sigma Chemical Co., St. Louis, MO) may be employed. The immobilized polypeptide is then incubated with the sample, and antibody is allowed to bind to the antigen. The sample may be diluted with a suitable diluent, such as phosphate-buffered saline (PBS) prior to incubation. In general, an appropriate contact time (*i.e.*, incubation time) is that period of time that is sufficient to detect the presence of antibody within a *M. tuberculosis*-infected sample. Preferably, the contact time is sufficient to achieve a level

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of binding that is at least 95% of that achieved at equilibrium between bound and unbound antibody. Those of ordinary skill in the art will recognize that the time necessary to achieve equilibrium may be readily determined by assaying the level of binding that occurs over a period of time. At room temperature, an incubation time of about 30 minutes is generally sufficient.

Unbound sample may then be removed by washing the solid support with an appropriate buffer, such as PBS containing 0.1% Tween 20TM. Detection reagent may then be added to the solid support. An appropriate detection reagent is any compound that binds to the immobilized antibody-polypeptide complex and that can be detected by any of a variety of means known to those in the art. Preferably, the detection reagent contains a binding agent (such as, for example, Protein A, Protein G, immunoglobulin, lectin or free antigen) conjugated to a reporter group. Preferred reporter groups include enzymes (such as horseradish peroxidase), substrates, cofactors, inhibitors, dyes, radionuclides, luminescent groups, fluorescent groups and biotin. The conjugation of binding agent to reporter group may be achieved using standard methods known to those of ordinary skill in the art. Common binding agents may also be purchased conjugated to a variety of reporter groups from many commercial sources (e.g., Zymed Laboratories, San Francisco, CA, and Pierce, Rockford, IL).

The detection reagent is then incubated with the immobilized antibody-polypeptide complex for an amount of time sufficient to detect the bound antibody. An appropriate amount of time may generally be determined from the manufacturer's instructions or by assaying the level of binding that occurs over a period of time. Unbound detection reagent is then removed and bound detection reagent is detected using the reporter group. The method employed for detecting the reporter group depends upon the nature of the reporter group. For radioactive groups, scintillation counting or autoradiographic methods are generally appropriate. Spectroscopic methods may be used to detect dyes, luminescent groups and fluorescent groups. Biotin may be detected using avidin, coupled to a different reporter group (commonly a radioactive or fluorescent group or an enzyme). Enzyme reporter groups may generally be detected by the addition of substrate (generally for a specific period of time), followed by spectroscopic or other analysis of the reaction products.

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To determine the presence or absence of anti-M. tuberculosis antibodies in the sample, the signal detected from the reporter group that remains bound to the solid support is generally compared to a signal that corresponds to a predetermined cut-off value. In one preferred embodiment, the cut-off value is the average mean signal obtained when the immobilized antigen is incubated with samples from an uninfected patient. In general, a sample generating a signal that is three standard deviations above the predetermined cut-off value is considered positive for tuberculosis. In an alternate preferred embodiment, the cutoff value is determined using a Receiver Operator Curve, according to the method of Sackett et al., Clinical Epidemiology: A Basic Science for Clinical Medicine, Little Brown and Co., 1985, pp. 106-107. Briefly, in this embodiment, the cut-off value may be determined from a plot of pairs of true positive rates (i.e., sensitivity) and false positive rates (100%-specificity) that correspond to each possible cut-off value for the diagnostic test result. The cut-off value on the plot that is the closest to the upper left-hand corner (i.e., the value that encloses the largest area) is the most accurate cut-off value, and a sample generating a signal that is higher than the cut-off value determined by this method may be considered positive. Alternatively, the cut-off value may be shifted to the left along the plot, to minimize the false positive rate, or to the right, to minimize the false negative rate. In general, a sample generating a signal that is higher than the cut-off value determined by this method is considered positive for tuberculosis.

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In a related embodiment, the assay is performed in a rapid flow-through or strip test format, wherein the antigen is immobilized on a membrane, such as nitrocellulose. In the flow-through test, antibodies within the sample bind to the immobilized polypeptide as the sample passes through the membrane. A detection reagent (e.g., protein A-colloidal gold) then binds to the antibody-polypeptide complex as the solution containing the detection reagent flows through the membrane. The detection of bound detection reagent may then be performed as described above. In the strip test format, one end of the membrane to which polypeptide is bound is immersed in a solution containing the sample. The sample migrates along the membrane through a region containing detection reagent and to the area of immobilized polypeptide. Concentration of detection reagent at the polypeptide indicates the presence of anti-M. tuberculosis antibodies in the sample. Typically, the concentration of

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detection reagent at that site generates a pattern, such as a line, that can be read visually. The absence of such a pattern indicates a negative result. In general, the amount of polypeptide immobilized on the membrane is selected to generate a visually discernible pattern when the biological sample contains a level of antibodies that would be sufficient to generate a positive signal in an ELISA, as discussed above. Preferably, the amount of polypeptide immobilized on the membrane ranges from about 25 ng to about 1 μ g, and more preferably from about 50 ng to about 500 ng. Such tests can typically be performed with a very small amount (e.g., one drop) of patient serum or blood.

Of course, numerous other assay protocols exist that are suitable for use with the polypeptides of the present invention. The above descriptions are intended to be exemplary only.

In yet another aspect, the present invention provides antibodies to the inventive polypeptides. Antibodies may be prepared by any of a variety of techniques known to those of ordinary skill in the art. See, e.g., Harlow and Lane, Antibodies: A Laboratory Manual, Cold Spring Harbor Laboratory, 1988. In one such technique, an immunogen comprising the antigenic polypeptide is initially injected into any of a wide variety of mammals (e.g., mice, rats, rabbits, sheep and goats). In this step, the polypeptides of this invention may serve as the immunogen without modification. Alternatively, particularly for relatively short polypeptides, a superior immune response may be elicited if the polypeptide is joined to a carrier protein, such as bovine serum albumin or keyhole limpet hemocyanin. The immunogen is injected into the animal host, preferably according to a predetermined schedule incorporating one or more booster immunizations, and the animals are bled periodically. Polyclonal antibodies specific for the polypeptide may then be purified from such antisera by, for example, affinity chromatography using the polypeptide coupled to a suitable solid support.

Monoclonal antibodies specific for the antigenic polypeptide of interest may be prepared, for example, using the technique of Kohler and Milstein, *Eur. J. Immunol.* 6:511-519, 1976, and improvements thereto. Briefly, these methods involve the preparation of immortal cell lines capable of producing antibodies having the desired specificity (*i.e.*, reactivity with the polypeptide of interest). Such cell lines may be produced, for example,

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from spleen cells obtained from an animal immunized as described above. The spleen cells are then immortalized by, for example, fusion with a myeloma cell fusion partner, preferably one that is syngeneic with the immunized animal. A variety of fusion techniques may be employed. For example, the spleen cells and myeloma cells may be combined with a nonionic detergent for a few minutes and then plated at low density on a selective medium that supports the growth of hybrid cells, but not myeloma cells. A preferred selection technique uses HAT (hypoxanthine, aminopterin, thymidine) selection. After a sufficient time, usually about 1 to 2 weeks, colonies of hybrids are observed. Single colonies are selected and tested for binding activity against the polypeptide. Hybridomas having high reactivity and specificity are preferred.

Monoclonal antibodies may be isolated from the supernatants of growing hybridoma colonies. In addition, various techniques may be employed to enhance the yield, such as injection of the hybridoma cell line into the peritoneal cavity of a suitable vertebrate host, such as a mouse. Monoclonal antibodies may then be harvested from the ascites fluid or the blood. Contaminants may be removed from the antibodies by conventional techniques, such as chromatography, gel filtration, precipitation, and extraction. The polypeptides of this invention may be used in the purification process in, for example, an affinity chromatography step.

Antibodies may be used in diagnostic tests to detect the presence of *M. tuberculosis* antigens using assays similar to those detailed above and other techniques well known to those of skill in the art, thereby providing a method for detecting *M. tuberculosis* infection in a patient.

Diagnostic reagents of the present invention may also comprise DNA sequences encoding one or more of the above polypeptides, or one or more portions thereof. For example, at least two oligonucleotide primers may be employed in a polymerase chain reaction (PCR) based assay to amplify *M. tuberculosis*-specific cDNA derived from a biological sample, wherein at least one of the oligonucleotide primers is specific for a DNA molecule encoding a polypeptide of the present invention. The presence of the amplified cDNA is then detected using techniques well known in the art, such as gel electrophoresis. Similarly, oligonucleotide probes specific for a DNA molecule encoding a polypeptide of the

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present invention may be used in a hybridization assay to detect the presence of an inventive polypeptide in a biological sample.

As used herein, the term "oligonucleotide primer/probe specific for a DNA molecule" means an oligonucleotide sequence that has at least about 80%, preferably at least about 90% and more preferably at least about 95%, identity to the DNA molecule in question. Oligonucleotide primers and/or probes which may be usefully employed in the inventive diagnostic methods preferably have at least about 10-40 nucleotides. In a preferred embodiment, the oligonucleotide primers comprise at least about 10 contiguous nucleotides of a DNA molecule encoding one of the polypeptides disclosed herein. oligonucleotide probes for use in the inventive diagnostic methods comprise at least about 15 contiguous oligonucleotides of a DNA molecule encoding one of the polypeptides disclosed herein. Techniques for both PCR based assays and hybridization assays are well known in the art (see, for example, Mullis et al. Ibid; Ehrlich, Ibid). Primers or probes may thus be used to detect M. tuberculosis-specific sequences in biological samples. DNA probes or primers comprising oligonucleotide sequences described above may be used alone, in combination with each other, or with previously identified sequences, such as the 38 kD antigen discussed above.

The following Examples are offered by way of illustration and not by way of limitation.

EXAMPLES

EXAMPLE 1

PURIFICATION AND CHARACTERIZATION OF POLYPEPTIDES
FROM M. TUBERCULOSIS CULTURE FILTRATE

This example illustrates the preparation of *M. tuberculosis* soluble polypeptides from culture filtrate. Unless otherwise noted, all percentages in the following example are weight per volume.

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M. tuberculosis (either H37Ra, ATCC No. 25177, or H37Rv, ATCC No. 25618) was cultured in sterile GAS media at 37°C for fourteen days. The media was then vacuum filtered (leaving the bulk of the cells) through a 0.45 μ filter into a sterile 2.5 L bottle. The media was then filtered through a 0.2 μ filter into a sterile 4 L bottle. NaN₃ was then added to the culture filtrate to a concentration of 0.04%. The bottles were then placed in a 4°C cold room.

The culture filtrate was concentrated by placing the filtrate in a 12 L reservoir that had been autoclaved and feeding the filtrate into a 400 ml Amicon stir cell which had been rinsed with ethanol and contained a 10,000 kDa MWCO membrane. The pressure was maintained at 60 psi using nitrogen gas. This procedure reduced the 12 L volume to approximately 50 ml.

The culture filtrate was then dialyzed into 0.1% ammonium bicarbonate using a 8,000 kDa MWCO cellulose ester membrane, with two changes of ammonium bicarbonate solution. Protein concentration was then determined by a commercially available BCA assay (Pierce, Rockford, IL).

The dialyzed culture filtrate was then lyophilized, and the polypeptides resuspended in distilled water. The polypeptides were then dialyzed against 0.01 mM 1,3 bis[tris(hydroxymethyl)-methylamino]propane, pH 7.5 (Bis-Tris propane buffer), the initial conditions for anion exchange chromatography. Fractionation was performed using gel profusion chromatography on a POROS 146 II Q/M anion exchange column 4.6 mm x 100 mm (Perseptive BioSystems, Framingham, MA) equilibrated in 0.01 mM Bis-Tris propane buffer pH 7.5. Polypeptides were eluted with a linear 0-0.5 M NaCl gradient in the above buffer system. The column eluent was monitored at a wavelength of 220 nm.

The pools of polypeptides eluting from the ion exchange column were dialyzed against distilled water and lyophilized. The resulting material was dissolved in 0.1% trifluoroacetic acid (TFA) pH 1.9 in water, and the polypeptides were purified on a Delta-Pak C18 column (Waters, Milford, MA) 300 Angstrom pore size, 5 micron particle size (3.9 x 150 mm). The polypeptides were eluted from the column with a linear gradient from 0-60% dilution buffer (0.1% TFA in acetonitrile). The flow rate was 0.75 ml/minute and the HPLC eluent was monitored at 214 nm. Fractions containing the eluted polypeptides were collected

to maximize the purity of the individual samples. Approximately 200 purified polypeptides were obtained.

The purified polypeptides were then screened for the ability to induce T-cell proliferation in PBMC preparations. The PBMCs from donors known to be PPD skin test positive and whose T cells were shown to proliferate in response to PPD and crude soluble proteins from MTB were cultured in medium comprising RPMI 1640 supplemented with 10% pooled human serum and 50 μ g/ml gentamicin. Purified polypeptides were added in duplicate at concentrations of 0.5 to 10 μ g/mL. After six days of culture in 96-well round-bottom plates in a volume of 200 μ l, 50 μ l of medium was removed from each well for determination of IFN- γ levels, as described below. The plates were then pulsed with 1 μ Ci/well of tritiated thymidine for a further 18 hours, harvested and tritium uptake determined using a gas scintillation counter. Fractions that resulted in proliferation in both replicates three fold greater than the proliferation observed in cells cultured in medium alone were considered positive.

IFN-γ was measured using an enzyme-linked immunosorbent assay (ELISA). ELISA plates were coated with a mouse monoclonal antibody directed to human IFN-γ (Chemicon) in PBS for four hours at room temperature. Wells were then blocked with PBS containing 5% (W/V) non-fat dried milk for 1 hour at room temperature. The plates were then washed six times in PBS/0.2% TWEEN-20 and samples diluted 1:2 in culture medium in the ELISA plates were incubated overnight at room temperature. The plates were again washed and a polyclonal rabbit anti-human IFN-γ serum diluted 1:3000 in PBS/10% normal goat serum was added to each well. The plates were then incubated for two hours at room temperature, washed and horseradish peroxidase-coupled anti-rabbit IgG (Jackson Labs.) was added at a 1:2000 dilution in PBS/5% non-fat dried milk. After a further two hour incubation at room temperature, the plates were washed and TMB substrate added. The reaction was stopped after 20 min with 1 N sulfuric acid. Optical density was determined at 450 nm using 570 nm as a reference wavelength. Fractions that resulted in both replicates giving an OD two fold greater than the mean OD from cells cultured in medium alone, plus 3 standard deviations, were considered positive.

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For sequencing, the polypeptides were individually dried onto BiobreneTM (Perkin Elmer/Applied BioSystems Division, Foster City, CA) treated glass fiber filters. The filters with polypeptide were loaded onto a Perkin Elmer/Applied BioSystems Division Procise 492 protein sequencer. The polypeptides were sequenced from the amino terminal and using traditional Edman chemistry. The amino acid sequence was determined for each polypeptide by comparing the retention time of the PTH amino acid derivative to the appropriate PTH derivative standards.

Using the procedure described above, antigens having the following N-terminal sequences were isolated:

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- (a) Asp-Pro-Val-Asp-Ala-Val-Ile-Asn-Thr-Thr-Xaa-Asn-Tyr-Gly-Gln-Val-Val-Ala-Ala-Leu (SEQ ID NO: 54);
- (b) Ala-Val-Glu-Ser-Gly-Met-Leu-Ala-Leu-Gly-Thr-Pro-Ala-Pro-Ser (SEQ ID NO: 55);
- (c) Ala-Ala-Met-Lys-Pro-Arg-Thr-Gly-Asp-Gly-Pro-Leu-Glu-Ala-Ala-Lys-Glu-Gly-Arg (SEQ ID NO: 56);

(d) Tyr-Tyr-Trp-Cys-Pro-Gly-Gln-Pro-Phe-Asp-Pro-Ala-Trp-Gly-Pro (SEO ID NO: 57);

(e) Asp-Ile-Gly-Ser-Glu-Ser-Thr-Glu-Asp-Gln-Gln-Xaa-Ala-Val (SEQ ID NO: 58);

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- (f) Ala-Glu-Glu-Ser-Ile-Ser-Thr-Xaa-Glu-Xaa-Ile-Val-Pro (SEQ ID NO: 59);
- (g) Asp-Pro-Glu-Pro-Ala-Pro-Pro-Val-Pro-Thr-Ala-Ala-Ala-Ala-Pro-Pro-Ala (SEQ ID NO: 60); and
- (h) Ala-Pro-Lys-Thr-Tyr-Xaa-Glu-Glu-Leu-Lys-Gly-Thr-Asp-Thr-Gly (SEQ ID NO: 61);

wherein Xaa may be any amino acid.

An additional antigen was isolated employing a microbore HPLC purification step in addition to the procedure described above. Specifically, 20 µl of a fraction comprising a mixture of antigens from the chromatographic purification step previously described, was purified on an Aquapore C18 column (Perkin Elmer/Applied Biosystems Division, Foster

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City, CA) with a 7 micron pore size, column size 1 mm x 100 mm, in a Perkin Elmer/Applied Biosystems Division Model 172 HPLC. Fractions were eluted from the column with a linear gradient of 1%/minute of acetonitrile (containing 0.05% TFA) in water (0.05% TFA) at a flow rate of 80 µl/minute. The eluent was monitored at 250 nm. The original fraction was separated into 4 major peaks plus other smaller components and a polypeptide was obtained which was shown to have a molecular weight of 12.054 Kd (by mass spectrometry) and the following N-terminal sequence:

 (i) Asp-Pro-Ala-Ser-Ala-Pro-Asp-Val-Pro-Thr-Ala-Ala-Gln-Gln-Thr-Ser-Leu-Leu-Asn-Asn-Leu-Ala-Asp-Pro-Asp-Val-Ser-Phe-Ala-Asp (SEQ ID NO: 62).

This polypeptide was shown to induce proliferation and IFN-γ production in PBMC preparations using the assays described above.

Additional soluble antigens were isolated from *M. tuberculosis* culture filtrate as follows. *M. tuberculosis* culture filtrate was prepared as described above. Following dialysis against Bis-Tris propane buffer, at pH 5.5, fractionation was performed using anion exchange chromatography on a Poros QE column 4.6 x 100 mm (Perseptive Biosystems) equilibrated in Bis-Tris propane buffer pH 5.5. Polypeptides were eluted with a linear 0-1.5 M NaCl gradient in the above buffer system at a flow rate of 10 ml/min. The column eluent was monitored at a wavelength of 214 nm.

The fractions eluting from the ion exchange column were pooled and subjected to reverse phase chromatography using a Poros R2 column 4.6 x 100 mm (Perseptive Biosystems). Polypeptides were eluted from the column with a linear gradient from 0-100% acetonitrile (0.1% TFA) at a flow rate of 5 ml/min. The eluent was monitored at 214 nm.

Fractions containing the eluted polypeptides were lyophilized and resuspended in 80 μl of aqueous 0.1% TFA and further subjected to reverse phase chromatography on a Vydac C4 column 4.6 x 150 mm (Western Analytical, Temecula, CA) with a linear gradient of 0-100% acetonitrile (0.1% TFA) at a flow rate of 2 ml/min. Eluent was monitored at 214 nm.

The fraction with biological activity was separated into one major peak plus other smaller components. Western blot of this peak onto PVDF membrane revealed three major bands of molecular weights 14 Kd, 20 Kd and 26 Kd. These polypeptides were determined to have the following N-terminal sequences, respectively:

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- (j) Xaa-Asp-Ser-Glu-Lys-Ser-Ala-Thr-Ile-Lys-Val-Thr-Asp-Ala-Ser; (SEQ ID NO: 129)
- (k) Ala-Gly-Asp-Thr-Xaa-Ile-Tyr-Ile-Val-Gly-Asn-Leu-Thr-Ala-Asp; (SEQ ID NO: 130) and
- (l) Ala-Pro-Glu-Ser-Gly-Ala-Gly-Leu-Gly-Gly-Thr-Val-Gln-Ala-Gly; (SEQ ID NO: 131), wherein Xaa may be any amino acid.

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Using the assays described above, these polypeptides were shown to induce proliferation and IFN- γ production in PBMC preparations. Figs. 1A and B show the results of such assays using PBMC preparations from a first and a second donor, respectively.

DNA sequences that encode the antigens designated as (a), (c), (d) and (g) above were obtained by screening a *M. tuberculosis* genomic library using ³²P end labeled degenerate oligonucleotides corresponding to the N-terminal sequence and containing *M. tuberculosis* codon bias. The screen performed using a probe corresponding to antigen (a) above identified a clone having the sequence provided in SEQ ID NO: 96. The polypeptide encoded by SEQ ID NO: 96 is provided in SEQ ID NO: 97. The screen performed using a probe corresponding to antigen (g) above identified a clone having the sequence provided in SEQ ID NO: 52. The polypeptide encoded by SEQ ID NO: 52 is provided in SEQ ID NO: 53. The screen performed using a probe corresponding to antigen (d) above identified a clone having the sequence provided in SEQ ID NO: 24, and the screen performed with a probe corresponding to antigen (c) identified a clone having the sequence provided in SEQ ID NO: 25.

The above amino acid sequences were compared to known amino acid sequences in the gene bank using the DNA STAR system. The database searched contains some 173,000 proteins and is a combination of the Swiss, PIR databases along with translated protein sequences (Version 87). No significant homologies to the amino acid sequences for antigens (a)-(h) and (l) were detected.

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The amino acid sequence for antigen (i) was found to be homologous to a sequence from *M. leprae*. The full length *M. leprae* sequence was amplified from genomic DNA using the sequence obtained from GENBANK. This sequence was then used to screen an *M. tuberculosis* library and a full length copy of the *M. tuberculosis* homologue was obtained (SEQ ID NO: 94).

The amino acid sequence for antigen (j) was found to be homologous to a known *M. tuberculosis* protein translated from a DNA sequence. To the best of the inventors' knowledge, this protein has not been previously shown to possess T-cell stimulatory activity. The amino acid sequence for antigen (k) was found to be related to a sequence from *M. leprae*.

In the proliferation and IFN-γ assays described above, using three PPD positive donors, the results for representative antigens provided above are presented in Table 1:

15 <u>TABLE 1</u>

RESULTS OF PBMC PROLIFERATION AND IFN-y ASSAYS

Sequence	Proliferation	IFN-γ
(a)	+	•
(c)	+++	+++
(d)	++	++
(g)	+++	+++
(h)	+++	+++

In Table 1, responses that gave a stimulation index (SI) of between 2 and 4 20 (compared to cells cultured in medium alone) were scored as +, as SI of 4-8 or 2-4 at a concentration of 1 μg or less was scored as ++ and an SI of greater than 8 was scored as +++. The antigen of sequence (i) was found to have a high SI (+++) for one donor and lower SI (++ and +) for the two other donors in both proliferation and IFN-γ assays. These results

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indicate that these antigens are capable of inducing proliferation and/or interferon-y production.

EXAMPLE 2

USE OF PATIENT SERA TO ISOLATE M. TUBERCULOSIS ANTIGENS

This example illustrates the isolation of antigens from M. tuberculosis lysate by screening with serum from M. tuberculosis-infected individuals.

Dessicated *M. tuberculosis* H37Ra (Difco Laboratories) was added to a 2% NP40 solution, and alternately homogenized and sonicated three times. The resulting suspension was centrifuged at 13,000 rpm in microfuge tubes and the supernatant put through a 0.2 micron syringe filter. The filtrate was bound to Macro Prep DEAE beads (BioRad, Hercules, CA). The beads were extensively washed with 20 mM Tris pH 7.5 and bound proteins eluted with 1M NaCl. The NaCl elute was dialyzed overnight against 10 mM Tris, pH 7.5. Dialyzed solution was treated with DNase and RNase at 0.05 mg/ml for 30 min. at room temperature and then with α-D-mannosidase, 0.5 U/mg at pH 4.5 for 3-4 hours at room temperature. After returning to pH 7.5, the material was fractionated via FPLC over a Bio Scale-Q-20 column (BioRad). Fractions were combined into nine pools, concentrated in a Centriprep 10 (Amicon, Beverley, MA) and screened by Western blot for serological activity using a serum pool from *M. tuberculosis*-infected patients which was not immunoreactive with other antigens of the present invention.

The most reactive fraction was run in SDS-PAGE and transferred to PVDF. A band at approximately 85 Kd was cut out yielding the sequence:

(m) Xaa-Tyr-Ile-Ala-Tyr-Xaa-Thr-Thr-Ala-Gly-Ile-Val-Pro-Gly-Lys-Ile-Asn-Val-His-Leu-Val; (SEQ ID NO: 132), wherein Xaa may be any amino acid.

Comparison of this sequence with those in the gene bank as described above, revealed no significant homologies to known sequences.

A DNA sequence that encodes the antigen designated as (m) above was obtained by screening a genomic M. tuberculosis Erdman strain library using labeled

degenerate oligonucleotides corresponding to the N-terminal sequence of SEQ ID NO:137. A clone was identified having the DNA sequence provided in SEQ ID NO: 198. This sequence was found to encode the amino acid sequence provided in SEQ ID NO: 199. Comparison of these sequences with those in the genebank revealed some similarity to sequences previously identified in *M. tuberculosis* and *M. bovis*.

EXAMPLE 3

PREPARATION OF DNA SEQUENCES ENCODING M. TUBERCULOSIS ANTIGENS

This example illustrates the preparation of DNA sequences encoding *M. tuberculosis* antigens by screening a *M. tuberculosis* expression library with sera obtained from patients infected with *M. tuberculosis*, or with anti-sera raised against *M. tuberculosis* antigens.

15 A. PREPARATION OF M. TUBERCULOSIS SOLUBLE ANTIGENS USING RABBIT ANTI-SERA RAISED AGAINST M. TUBERCULOSIS SUPERNATANT

Genomic DNA was isolated from the *M. tuberculosis* strain H37Ra. The DNA was randomly sheared and used to construct an expression library using the Lambda ZAP expression system (Stratagene, La Jolla, CA). Rabbit anti-sera was generated against secretory proteins of the *M. tuberculosis* strains H37Ra, H37Rv and Erdman by immunizing a rabbit with concentrated supernatant of the *M. tuberculosis* cultures. Specifically, the rabbit was first immunized subcutaneously with 200 µg of protein antigen in a total volume of 2 ml containing 100 µg muramyl dipeptide (Calbiochem, La Jolla, CA) and 1 ml of incomplete Freund's adjuvant. Four weeks later the rabbit was boosted subcutaneously with 100 µg antigen in incomplete Freund's adjuvant. Finally, the rabbit was immunized intravenously four weeks later with 50 µg protein antigen. The anti-sera were used to screen the expression library as described in Sambrook et al., *Molecular Cloning: A Laboratory Manual*, Cold Spring Harbor Laboratories, Cold Spring Harbor, NY, 1989. Bacteriophage plaques expressing immunoreactive antigens were purified. Phagemid from the plaques was rescued and the nucleotide sequences of the *M. tuberculosis* clones deduced.

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Thirty two clones were purified. Of these, 25 represent sequences that have not been previously identified in *M. tuberculosis*. Proteins were induced by IPTG and purified by gel elution, as described in Skeiky et al., *J. Exp. Med. 181*:1527-1537, 1995. Representative partial sequences of DNA molecules identified in this screen are provided in SEQ ID NOS: 1-25. The corresponding predicted amino acid sequences are shown in SEQ ID NOS: 64-88.

On comparison of these sequences with known sequences in the gene bank using the databases described above, it was found that the clones referred to hereinafter as TbRA2A, TbRA16, TbRA18, and TbRA29 (SEQ ID NOS: 77, 69, 71, 76) show some homology to sequences previously identified in *Mycobacterium leprae* but not in *M. tuberculosis*. TbRA11, TbRA26, TbRA28 and TbDPEP (SEQ ID NOS: 66, 74, 75, 53) have been previously identified in *M. tuberculosis*. No significant homologies were found to TbRA1, TbRA3, TbRA4, TbRA9, TbRA10, TbRA13, TbRA17, TbRA19, TbRA29, TbRA32, TbRA36 and the overlapping clones TbRA35 and TbRA12 (SEQ ID NOS: 64, 78, 82, 83, 65, 68, 76, 72, 76, 79, 81, 80, 67, respectively). The clone TbRa24 is overlapping with clone TbRa29.

B. <u>Use of Sera from Patients Having Pulmonary or Pleural Tuberculosis to</u> IDENTIFY DNA Sequences Encoding M. Tuberculosis Antigens

The genomic DNA library described above, and an additional H37Rv library, were screened using pools of sera obtained from patients with active tuberculosis. To prepare the H37Rv library, *M. tuberculosis* strain H37Rv genomic DNA was isolated, subjected to partial Sau3A digestion and used to construct an expression library using the Lambda Zap expression system (Stratagene, La Jolla, Ca). Three different pools of sera, each containing sera obtained from three individuals with active pulmonary or pleural disease, were used in the expression screening. The pools were designated TbL, TbM and TbH, referring to relative reactivity with H37Ra lysate (*i.e.*, TbL = low reactivity, TbM = medium reactivity and TbH = high reactivity) in both ELISA and immunoblot format. A fourth pool of sera from seven patients with active pulmonary tuberculosis was also employed. All of the sera

lacked increased reactivity with the recombinant 38 kD M. tuberculosis H37Ra phosphate-binding protein.

All pools were pre-adsorbed with *E. coli* lysate and used to screen the H37Ra and H37Rv expression libraries, as described in Sambrook et al., *Molecular Cloning: A Laboratory Manual*, Cold Spring Harbor Laboratories, Cold Spring Harbor, NY, 1989. Bacteriophage plaques expressing immunoreactive antigens were purified. Phagemid from the plaques was rescued and the nucleotide sequences of the *M. tuberculosis* clones deduced.

Thirty two clones were purified. Of these, 31 represented sequences that had not been previously identified in human *M. tuberculosis*. Representative sequences of the DNA molecules identified are provided in SEQ ID NOS:: 26-51 and 100. Of these, TbH-8-2 (SEQ. ID NO. 100) is a partial clone of TbH-8, and TbH-4 (SEQ. ID NO. 43) and TbH-4-FWD (SEQ. ID NO. 44) are non-contiguous sequences from the same clone. Amino acid sequences for the antigens hereinafter identified as Tb38-1, TbH-4, TbH-8, TbH-9, and TbH-12 are shown in SEQ ID NOS.: 89-93. Comparison of these sequences with known sequences in the gene bank using the databases identified above revealed no significant homologies to TbH-4, TbH-8, TbH-9 and TbM-3, although weak homologies were found to TbH-9. TbH-12 was found to be homologous to a 34 kD antigenic protein previously identified in *M. paratuberculosis* (Acc. No. S28515). Tb38-1 was found to be located 34 base pairs upstream of the open reading frame for the antigen ESAT-6 previously identified in *M. bovis* (Acc. No. U34848) and in *M. tuberculosis* (Sorensen et al., *Infec. Immun.* 63:1710-1717, 1995).

Probes derived from Tb38-1 and TbH-9, both isolated from an H37Ra library, were used to identify clones in an H37Rv library. Tb38-1 hybridized to Tb38-1F2, Tb38-1F3, Tb38-1F5 and Tb38-1F6 (SEQ. ID NOS: 107, 108, 111, 113, and 114). (SEQ ID NOS: 107 and 108 are non-contiguous sequences from clone Tb38-1F2.) Two open reading frames were deduced in Tb38-IF2; one corresponds to Tb37FL (SEQ. ID. NO. 109), the second, a partial sequence, may be the homologue of Tb38-1 and is called Tb38-IN (SEQ. ID NO. 110). The deduced amino acid sequence of Tb38-1F3 is presented in SEQ. ID. NO. 112. A TbH-9 probe identified three clones in the H37Rv library: TbH-9-FL (SEQ. ID NO. 101), which may be the homologue of TbH-9 (R37Ra), TbH-9-1 (SEQ. ID NO. 103), and TbH-8-2 (SEQ.

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ID NO. 105) is a partial clone of TbH-8. The deduced amino acid sequences for these three clones are presented in SEQ ID NOS: 102, 104 and 106.

Further screening of the *M. tuberculosis* genomic DNA library, as described above, resulted in the recovery of ten additional reactive clones, representing seven different genes. One of these genes was identified as the 38 Kd antigen discussed above, one was determined to be identical to the 14Kd alpha crystallin heat shock protein previously shown to be present in *M. tuberculosis*, and a third was determined to be identical to the antigen TbH-8 described above. The determined DNA sequences for the remaining five clones (hereinafter referred to as TbH-29, TbH-30, TbH-32 and TbH-33) are provided in SEQ ID NO: 133-136, respectively, with the corresponding predicted amino acid sequences being provided in SEQ ID NO: 137-140, respectively. The DNA and amino acid sequences for these antigens were compared with those in the gene bank as described above. No homologies were found to the 5' end of TbH-29 (which contains the reactive open reading frame), although the 3' end of TbH-29 was found to be identical to the *M. tuberculosis* cosmid Y227. TbH-32 and TbH-33 were found to be identical to the previously identified *M. tuberculosis* insertion element IS6110 and to the *M. tuberculosis* cosmid Y50, respectively. No significant homologies to TbH-30 were found.

Positive phagemid from this additional screening were used to infect *E. coli* XL-1 Blue MRF', as described in Sambrook et al., *supra*. Induction of recombinant protein was accomplished by the addition of IPTG. Induced and uninduced lysates were run in duplicate on SDS-PAGE and transferred to nitrocellulose filters. Filters were reacted with human *M. tuberculosis* sera (1:200 dilution) reactive with TbH and a rabbit sera (1:200 or 1:250 dilution) reactive with the N-terminal 4 Kd portion of lacZ. Sera incubations were performed for 2 hours at room temperature. Bound antibody was detected by addition of ¹²⁵I-labeled Protein A and subsequent exposure to film for variable times ranging from 16 hours to 11 days. The results of the immunoblots are summarized in Table 2.

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TABLE 2

5	Antigen	Human M. tb <u>Sera</u>	Anti-lacZ <u>Sera</u>
	Тън-29	45 Kd	45 Kd
	ТЬН-30	No reactivity	29 Kd
	ТьН-32	12 Kd	12 Kd
	ТЬН-33	16 Kd	16 Kd

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Positive reaction of the recombinant human *M. tuberculosis* antigens with both the human *M. tuberculosis* sera and anti-lacZ sera indicate that reactivity of the human *M. tuberculosis* sera is directed towards the fusion protein. Antigens reactive with the anti-lacZ sera but not with the human *M. tuberculosis* sera may be the result of the human *M. tuberculosis* sera recognizing conformational epitopes, or the antigen-antibody binding kinetics may be such that the 2 hour sera exposure in the immunoblot is not sufficient.

Studies were undertaken to determine whether the antigens TbH-9 and Tb38-1 represent cellular proteins or are secreted into *M. tuberculosis* culture media. In the first study, rabbit sera were raised against A) secretory proteins of *M. tuberculosis*, B) the known secretory recombinant *M. tuberculosis* antigen 85b, C) recombinant Tb38-1 and D) recombinant TbH-9, using protocols substantially as described in Example 3A. Total *M. tuberculosis* lysate, concentrated supernatant of *M. tuberculosis* cultures and the recombinant antigens 85b, TbH-9 and Tb38-1 were resolved on denaturing gels, immobilized on nitrocellulose membranes and duplicate blots were probed using the rabbit sera described above.

The results of this analysis using control sera (panel I) and antisera (panel II) against secretory proteins, recombinant 85b, recombinant Tb38-1 and recombinant TbH-9 are shown in Figures 2A-D, respectively, wherein the lane designations are as follows: 1) molecular weight protein standards; 2) 5 µg of *M. tuberculosis* lysate; 3) 5 µg secretory proteins; 4) 50 ng recombinant Tb38-1; 5) 50 ng recombinant TbH-9; and 6) 50 ng recombinant 85b. The recombinant antigens were engineered with six terminal histidine

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residues and would therefore be expected to migrate with a mobility approximately 1 kD larger that the native protein. In Figure 2D, recombinant TbH-9 is lacking approximately 10 kD of the full-length 42 kD antigen, hence the significant difference in the size of the immunoreactive native TbH-9 antigen in the lysate lane (indicated by an arrow). These results demonstrate that Tb38-1 and TbH-9 are intracellular antigens and are not actively secreted by *M. tuberculosis*.

The finding that TbH-9 is an intracellular antigen was confirmed by determining the reactivity of TbH-9-specific human T cell clones to recombinant TbH-9, secretory *M. tuberculosis* proteins and PPD. A TbH-9-specific T cell clone (designated 131TbH-9) was generated from PBMC of a healthy PPD-positive donor. The proliferative response of 131TbH-9 to secretory proteins, recombinant TbH-9 and a control *M. tuberculosis* antigen, TbRa11, was determined by measuring uptake of tritiated thymidine, as described in Example 1. As shown in Figure 3A, the clone 131TbH-9 responds specifically to TbH-9, showing that TbH-9 is not a significant component of *M. tuberculosis* secretory proteins. Figure 3B shows the production of IFN-γ by a second TbH-9-specific T cell clone (designated PPD 800-10) prepared from PBMC from a healthy PPD-positive donor, following stimulation of the T cell clone with secretory proteins, PPD or recombinant TbH-9. These results further confirm that TbH-9 is not secreted by *M. tuberculosis*.

20 C. USE OF SERA FROM PATIENTS HAVING EXTRAPULMONARY TUBERCULOSIS TO IDENTIFY DNA SEQUENCES ENCODING M. TUBERCULOSIS ANTIGENS

Genomic DNA was isolated from M. tuberculosis Erdman strain, randomly sheared and used to construct an expression library employing the Lambda ZAP expression system (Stratagene, La Jolla, CA). The resulting library was screened using pools of sera obtained from individuals with extrapulmonary tuberculosis, as described above in Example 3B, with the secondary antibody being goat anti-human IgG + A + M (H+L) conjugated with alkaline phosphatase.

Eighteen clones were purified. Of these, 4 clones (hereinafter referred to as XP14, XP24, XP31 and XP32) were found to bear some similarity to known sequences. The determined DNA sequences for XP14, XP24 and XP31 are provided in SEQ ID NOS: 151-

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153, respectively, with the 5' and 3' DNA sequences for XP32 being provided in SEQ ID NOS: 154 and 155, respectively. The predicted amino acid sequence for XP14 is provided in SEQ ID NO: 156. The reverse complement of XP14 was found to encode the amino acid sequence provided in SEQ ID NO: 157.

Comparison of the sequences for the remaining 14 clones (hereinafter referred to as XP1-XP6, XP17-XP19, XP22, XP25, XP27, XP30 and XP36) with those in the genebank as described above, revealed no homologies with the exception of the 3' ends of XP2 and XP6 which were found to bear some homology to known M. tuberculosis cosmids. The DNA sequences for XP27 and XP36 are shown in SEQ ID NOS: 158 and 159, respectively, with the 5' sequences for XP4, XP5, XP17 and XP30 being shown in SEQ ID NOS: 160-163, respectively, and the 5' and 3' sequences for XP2, XP3, XP6, XP18, XP19, XP22 and XP25 being shown in SEQ ID NOS: 164 and 165; 166 and 167; 168 and 169; 170 and 171; 172 and 173; 174 and 175; and 176 and 177, respectively. XP1 was found to overlap with the DNA sequences for TbH4, disclosed above. The full-length DNA sequence 15 -- for TbH4-XP1 is provided in SEQ ID NO: 178. This DNA sequence was found to contain an open reading frame encoding the amino acid sequence shown in SEQ ID NO: 179. The reverse complement of TbH4-XP1 was found to contain an open reading frame encoding the amino acid sequence shown in SEQ ID NO: 180. The DNA sequence for XP36 was found to contain two open reading frames encoding the amino acid sequence shown in SEQ ID NOS: 181 and 182, with the reverse complement containing an open reading frame encoding the amino acid sequence shown in SEQ ID NO: 183.

Recombinant XP1 protein was prepared as described above in Example 3B, with a metal ion affinity chromatography column being employed for purification. Recombinant XP1 was found to stimulate cell proliferation and IFN-γ production in T cells isolated from an M. tuberculosis-immune donors.

PREPARATION OF M. TUBERCULOSIS SOLUBLE ANTIGENS USING RABBIT ANTI-SERA <u>D.</u> RAISED AGAINST M. TUBERCULOSIS FRACTIONATED PROTEINS

M. tuberculosis lysate was prepared as described above in Example 2. The resulting material was fractionated by HPLC and the fractions screened by Western blot for 30

serological activity with a serum pool from *M. tuberculosis*-infected patients which showed little or no immunoreactivity with other antigens of the present invention. Rabbit anti-sera was generated against the most reactive fraction using the method described in Example 3A. The anti-sera was used to screen an *M. tuberculosis* Erdman strain genomic DNA expression library prepared as described above. Bacteriophage plaques expressing immunoreactive antigens were purified. Phagemid from the plaques was rescued and the nucleotide sequences of the *M. tuberculosis* clones determined.

Ten different clones were purified. Of these, one was found to be TbRa35, described above, and one was found to be the previously identified *M. tuberculosis* antigen, HSP60. Of the remaining eight clones, six (hereinafter referred to as RDIF2, RDIF5, RDIF8, RDIF10, RDIF11 and RDIF12) were found to bear some similarity to previously identified *M. tuberculosis* sequences. The determined DNA sequences for RDIF2, RDIF5, RDIF8, RDIF10 and RDIF11 are provided in SEQ ID NOS: 184-188, respectively, with the corresponding predicted amino acid sequences being provided in SEQ ID NOS: 189-193, respectively. The 5' and 3' DNA sequences for RDIF12 are provided in SEQ ID NOS: 194 and 195, respectively. No significant homologies were found to the antigen RDIF-7. The determined DNA and predicted amino acid sequences for RDIF7 are provided in SEQ ID NOS: 196 and 197, respectively. One additional clone, referred to as RDIF6 was isolated, however, this was found to be identical to RDIF5.

Recombinant RDIF6, RDIF8, RDIF10 and RDIF11 were prepared as described above. These antigens were found to stimulate cell proliferation and IFN- γ production in T cells isolated from *M. tuberculosis*-immune donors.

EXAMPLE 4

PURIFICATION AND CHARACTERIZATION OF A POLYPEPTIDE FROM TUBERCULIN PURIFIED PROTEIN DERIVATIVE

An M. tuberculosis polypeptide was isolated from tuberculin purified protein derivative (PPD) as follows.

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PPD was prepared as published with some modification (Seibert, F. et al., Tuberculin purified protein derivative. Preparation and analyses of a large quantity for standard. The American Review of Tuberculosis 44:9-25, 1941). *M. tuberculosis* Rv strain was grown for 6 weeks in synthetic medium in roller bottles at 37°C. Bottles containing the bacterial growth were then heated to 100°C in water vapor for 3 hours. Cultures were sterile filtered using a 0.22 μ filter and the liquid phase was concentrated 20 times using a 3 kD cutoff membrane. Proteins were precipitated once with 50% ammonium sulfate solution and eight times with 25% ammonium sulfate solution. The resulting proteins (PPD) were fractionated by reverse phase liquid chromatography (RP-HPLC) using a C18 column (7.8 x 300 mM; Waters, Milford, MA) in a Biocad HPLC system (Perseptive Biosystems, Framingham, MA). Fractions were eluted from the column with a linear gradient from 0-100% buffer (0.1% TFA in acetonitrile). The flow rate was 10 ml/minute and eluent was monitored at 214 nm and 280 nm.

Six fractions were collected, dried, suspended in PBS and tested individually in *M. tuberculosis*-infected guinea pigs for induction of delayed type hypersensitivity (DTH) reaction. One fraction was found to induce a strong DTH reaction and was subsequently fractionated further by RP-HPLC on a microbore Vydac C18 column (Cat. No. 218TP5115) in a Perkin Elmer/Applied Biosystems Division Model 172 HPLC. Fractions were eluted with a linear gradient from 5-100% buffer (0.05% TFA in acetonitrile) with a flow rate of 80 µl/minute. Eluent was monitored at 215 nm. Eight fractions were collected and tested for induction of DTH in *M. tuberculosis*-infected guinea pigs. One fraction was found to induce strong DTH of about 16 mm induration. The other fractions did not induce detectable DTH. The positive fraction was submitted to SDS-PAGE gel electrophoresis and found to contain a single protein band of approximately 12 kD molecular weight.

This polypeptide, herein after referred to as DPPD, was sequenced from the amino terminal using a Perkin Elmer/Applied Biosystems Division Procise 492 protein sequencer as described above and found to have the N-terminal sequence shown in SEQ ID NO:: 124. Comparison of this sequence with known sequences in the gene bank as described above revealed no known homologies. Four cyanogen bromide fragments of DPPD were isolated and found to have the sequences shown in SEQ ID NOS: 125-128.

EXAMPLE 5

SYNTHESIS OF SYNTHETIC POLYPEPTIDES

Polypeptides may be synthesized on a Millipore 9050 peptide synthesizer using FMOC chemistry with HPTU (O-Benzotriazole-N,N,N',N'-tetramethyluronium hexafluorophosphate) activation. A Gly-Cys-Gly sequence may be attached to the amino terminus of the peptide to provide a method of conjugation or labeling of the peptide. Cleavage of the peptides from the solid support may be carried out using the following cleavage mixture: trifluoroacetic acid:ethanedithiol:thioanisole:water:phenol (40:1:2:2:3). After cleaving for 2 hours, the peptides may be precipitated in cold methyl-t-butyl-ether. The peptide pellets may then be dissolved in water containing 0.1% trifluoroacetic acid (TFA) and lyophilized prior to purification by C18 reverse phase HPLC. A gradient of 0-60% acetonitrile (containing 0.1% TFA) in water (containing 0.1% TFA) may be used to elute the peptides. Following lyophilization of the pure fractions, the peptides may be characterized using electrospray mass spectrometry and by amino acid analysis.

This procedure was used to synthesize a TbM-1 peptide that contains one and a half repeats of a TbM-1 sequence. The TbM-1 peptide has the sequence GCGDRSGGNLDQIRLRRDRSGGNL (SEQ ID NO: 63).

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EXAMPLE 6

USE OF REPRESENTATIVE ANTIGENS FOR SERODIAGNOSIS OF TUBERCULOSIS

This Example illustrates the diagnostic properties of several representative antigens.

Assays were performed in 96-well plates were coated with 200 ng antigen diluted to $50~\mu L$ in carbonate coating buffer, pH 9.6. The wells were coated overnight at 4°C (or 2 hours at 37°C). The plate contents were then removed and the wells were blocked for 2 hours with 200 μL of PBS/1% BSA. After the blocking step, the wells were washed five

times with PBS/0.1% Tween 20™. 50 µL sera, diluted 1:100 in PBS/0.1% Tween 20™/0.1% BSA, was then added to each well and incubated for 30 minutes at room temperature. The plates were then washed again five times with PBS/0.1% Tween 20™.

The enzyme conjugate (horseradish peroxidase - Protein A, Zymed, San Francisco, CA) was then diluted 1:10,000 in PBS/0.1% Tween 20™/0.1% BSA, and 50 µL of the diluted conjugate was added to each well and incubated for 30 minutes at room temperature. Following incubation, the wells were washed five times with PBS/0.1% Tween 20^{TM} . $100 \,\mu\text{L}$ of tetramethylbenzidine peroxidase (TMB) substrate (Kirkegaard and Perry Laboratories, Gaithersburg, MD) was added, undiluted, and incubated for about 15 minutes. The reaction was stopped with the addition of 100 µL of 1 N H₂SO₄ to each well, and the plates were read at 450 nm.

Figure 4 shows the ELISA reactivity of two recombinant antigens isolated using method A in Example 3 (TbRa3 and TbRa9) with sera from M. tuberculosis positive and negative patients. The reactivity of these antigens is compared to that of bacterial lysate isolated from M. tuberculosis strain H37Ra (Difco, Detroit, MI). In both cases, the recombinant antigens differentiated positive from negative sera. Based on cut-off values obtained from receiver-operator curves, TbRa3 detected 56 out of 87 positive sera, and TbRa9 detected 111 out of 165 positive sera.

Figure 5 illustrates the ELISA reactivity of representative antigens isolated using method B of Example 3. The reactivity of the recombinant antigens TbH4, TbH12, Tb38-1 and the peptide TbM-1 (as described in Example 4) is compared to that of the 38 kD antigen described by Andersen and Hansen, Infect. Immun. 57:2481-2488, 1989. Again, all of the polypeptides tested differentiated positive from negative sera. Based on cut-off values obtained from receiver-operator curves, TbH4 detected 67 out of 126 positive sera, TbH12 detected 50 out of 125 positive sera, 38-1 detected 61 out of 101 positive sera and the TbM-1 25 peptide detected 25 out of 30 positive sera.

The reactivity of four antigens (TbRa3, TbRa9, TbH4 and TbH12) with sera from a group of M. tuberculosis infected patients with differing reactivity in the acid fast stain of sputum (Smithwick and David, Tubercle 52:226, 1971) was also examined, and compared

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to the reactivity of *M. tuberculosis* lysate and the 38 kD antigen. The results are presented in Table 3, below:

TABLE 3

REACTIVITY OF ANTIGENS WITH SERA FROM M. TUBERCULOSIS PATIENTS

	Acid Fast			ELISA	Values		
Patient	Sputum	Lysate	38kD	TbRa9	TbH12	TbH4	TbRa3
Tb01B93I-2	++++	1.853	0.634	0.998	1.022	1.030	# 1.314
Tb01B93I-19	++++	2.657	2.322	0.608	0.837	1.857	2.335
Tb01B93I-8	+++	2.703	0.527	0.492	0.281	0.501	2.002
Tb01B93I-10	+++	1.665	1.301	0.685	0.216	0.448	0.458
Tb01B93I-11	+++	2.817	0.697	0.509	0.301	0.173	2.608
Tb01B93I-15	+++	1.28	0.283	0.808	0.218	1.537	0.811
Tb01B93I-16	+++	2.908	>3	0.899	0.441	0.593	1.080
Tb01B93I-25	+++	0.395	0.131	0.335	0.211	0.107	0.948
Tb01B93I-87	+++	2.653	2.432	2.282	0.977	1.221	0.857
Tb01B93I-89	+++	1.912	2.370	2.436	0.876	0.520	0.952
Tb01B94I-108	+++	1.639	0.341	0.797	0.368	0.654	0.798
Tb01B94I-201	+++	1.721	0.419	0.661	0.137	0.064	0.692
Tb01B93I-88	++	1.939	1.269	2.519	1.381	0.214	0.530
Tb01B93I-92	++	2.355	2.329	2.78	0.685	0.997	2.527
Tb01B94I-109	++	0.993	0.620	0.574	0.441	0.5	2.558
Tb01B94I-210	++	2.777	>3	0.393	0.367	1.004	1.315
Tb01B94I-224	++	2.913	0.476	0.251	1.297	1.990	0.256

	Acid Fast			ELISA	Values		
Patient	Sputum	Lysate	38kD	TbRa9	ТьН12	ТЬН4	TbRa3
Tb01B93I-9	+	2.649	0.278	0.210	0.140	0.181	1.586
Tb01B93I-14	+	>3	1.538	0.282	0.291	0.549	2.880
Tb01B93I-21	+	2.645	0.739	2.499	0.783	0.536	1.770
Tb01B93I-22	+	0.714	0.451	2.082	0.285	0.269	1.159
Ть01В93І-31	+	0.956	0.490	1.019	0.812	0.176	1.293
Tb01B93I-32	_	2.261	0.786	0.668	0.273	0.535	0.405
Tb01B93I-52	_	0.658	0.114	0.434	0.330	0.273	1.140
Tb01B93I-99	-	2.118	0.584	1.62	0.119	0.977	0.729
Tb01B94I-130	-	1.349	0.224	0.86	0.282	0.383	2.146
Tb01B94I-131	-	0.685	0.324	1.173	0.059	0.118	1.431
AT4-0070	Normal	0.072	0.043	0.092	0.071	0.040	0.039
AT4-0105	Normal	0.397	0.121	0.118	0.103	0.078	0.390
3/15/94-1	Normal	0.227	0.064	0.098	0.026	0.001	0.228
4/15/93-2	Normal	0.114	0.240	0.071	0.034	0.041	0.264
5/26/94-4	Normal	0.089	0.259	0.096	0.046	0.008	0.053
5/26/94-3	Normal	0.139	0.093	0.085	0.019	0.067	0.01

Based on cut-off values obtained from receiver-operator curves, TbRa3 detected 23 out of 27 positive sera, TbRa9 detected 22 out of 27, TbH4 detected 18 out of 27 and TbH12 detected 15 out of 27. If used in combination, these four antigens would have a theoretical sensitivity of 27 out of 27, indicating that these antigens should complement each other in the serological detection of *M. tuberculosis* infection. In addition, several of the recombinant antigens detected positive sera that were not detected using the 38 kD antigen, indicating that these antigens may be complementary to the 38 kD antigen.

The reactivity of the recombinant antigen TbRa11 with sera from M. tuberculosis patients shown to be negative for the 38 kD antigen, as well as with sera from PPD positive and normal donors, was determined by ELISA as described above. The results are shown in Figure 6 which indicates that TbRa11, while being negative with sera from PPD positive and normal donors, detected sera that were negative with the 38 kD antigen. Of the thirteen 38 kD negative sera tested, nine were positive with TbRa11, indicating that this antigen may be reacting with a sub-group of 38 kD antigen negative sera. In contrast, in a group of 38 kD positive sera where TbRa11 was reactive, the mean OD 450 for TbRa11 was lower than that for the 38 kD antigen. The data indicate an inverse relationship between the presence of TbRa11 activity and 38 kD positivity.

The antigen TbRa2A was tested in an indirect ELISA using initially 50 µl of serum at 1:100 dilution for 30 minutes at room temperature followed by washing in PBS Tween and incubating for 30 minutes with biotinylated Protein A (Zymed, San Francisco, CA) at a 1:10,000 dilution. Following washing, 50 µl of streptavidin-horseradish peroxidase (Zymed) at 1:10,000 dilution was added and the mixture incubated for 30 minutes. After washing, the assay was developed with TMB substrate as described above. The reactivity of TbRa2A with sera from *M. tuberculosis* patients and normal donors in shown in Table 4. The mean value for reactivity of TbRa2A with sera from *M. tuberculosis* patients was 0.444 with a standard deviation of 0.309. The mean for reactivity with sera from normal donors was 0.109 with a standard deviation of 0.029. Testing of 38 kD negative sera (Figure 7) also indicated that the TbRa2A antigen was capable of detecting sera in this category.

TABLE 4

REACTIVITY OF TBRA2A WITH SERA FROM M. TUBERCULOSIS PATIENTS AND FROM NORMAL

DONORS

Serum ID	Status	OD 450
Tb85	TB	0.680
Tb86	TB	0.450
Tb87	TB	0.263
Tb88	TB	0.275
Tb89	TB	0.403

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Ть91	TB	0.393
Tb92	TB	0.401
Tb93	TB	0.232
Ть94	TB	0.333
Ть95	TB	0.435
Tb96	TB	0.284
Tb97	TB	0.320
Ть99	TB	0.328
Tb100	TB	0.817
Tb101	TB	0.607
Тъ102	TB	0.191
Tb103	TB	0.228
Tb107	TB	0.324
Tb109	TB	1.572
Tb112	TB	0.338
DL4-0176	Normal	0.036
AT4-0043	Normal	0.126
AT4-0044	Normal	0.130
AT4-0052	Normal	0.135
AT4-0053	Normal	0.133
AT4-0062	Normal	0.128
AT4-0070	Normal	0.088
AT4-0091	Normal	0.108
AT4-0100	Normal	0.106
AT4-0105	Normal	0.108
AT4-0109	Normal	0.105

The reactivity of the recombinant antigen (g) (SEQ ID NO: 60) with sera from *M. tuberculosis* patients and normal donors was determined by ELISA as described above. Figure 8 shows the results of the titration of antigen (g) with four *M. tuberculosis* positive sera that were all reactive with the 38 kD antigen and with four donor sera. All four positive sera were reactive with antigen (g).

The reactivity of the recombinant antigen TbH-29 (SEQ ID NO: 137) with sera from *M. tuberculosis* patients, PPD positive donors and normal donors was determined by indirect ELISA as described above. The results are shown in Figure 9. TbH-29 detected 30 out of 60 *M. tuberculosis* sera, 2 out of 8 PPD positive sera and 2 out of 27 normal sera.

Figure 10 shows the results of ELISA tests (both direct and indirect) of the antigen TbH-33 (SEQ ID NO: 140) with sera from *M. tuberculosis* patients and from normal

donors and with a pool of sera from *M. tuberculosis* patients. The mean OD 450 was demonstrated to be higher with sera from *M. tuberculosis* patients than from normal donors, with the mean OD 450 being significantly higher in the indirect ELISA than in the direct ELISA. Figure 11 is a titration curve for the reactivity of recombinant TbH-33 with sera from *M. tuberculosis* patients and from normal donors showing an increase in OD 450 with increasing concentration of antigen.

The reactivity of the recombinant antigens RDIF6, RDIF8 and RDIF10 (SEQ ID NOS: 184-187, respectively) with sera from *M. tuberculosis* patients and normal donors was determined by ELISA as described above. RDIF6 detected 6 out of 32 *M. tuberculosis* sera and 0 out of 15 normal sera; RDIF8 detected 14 out of 32 *M. tuberculosis* sera and 0 out of 15 normal sera; and RDIF10 detected 4 out of 27 *M. tuberculosis* sera and 1 out of 15 normal sera. In addition, RDIF10 was found to detect 0 out of 5 sera from PPD-positive donors.

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EXAMPLE 7

PREPARATION AND CHARACTERIZATION OF M. TUBERCULOSIS FUSION PROTEINS

A fusion protein containing TbRa3, the 38 kD antigen and Tb38-1 was prepared as follows.

Each of the DNA constructs TbRa3, 38 kD and Tb38-1 were modified by PCR in order to facilitate their fusion and the subsequent expression of the fusion protein TbRa3-38 kD-Tb38-1. TbRa3, 38 kD and Tb38-1 DNA was used to perform PCR using the primers PDM-64 and PDM-65 (SEQ ID NO: 141 and 142), PDM-57 and PDM-58 (SEQ ID NO: 143 and 144), and PDM-69 and PDM-60 (SEQ ID NO: 145-146), respectively. In each case, the DNA amplification was performed using 10 μl 10X Pfu buffer, 2 μl 10 mM dNTPs, 2 μl each of the PCR primers at 10 μM concentration, 81.5 μl water, 1.5 μl Pfu DNA polymerase (Stratagene, La Jolla, CA) and 1 μl DNA at either 70 ng/μl (for TbRa3) or 50 ng/μl (for 38 kD and Tb38-1). For TbRa3, denaturation at 94°C was performed for 2 min, followed by 40 cycles of 96°C for 15 sec and 72°C for 1 min, and lastly by 72°C for 4 min. For 38 kD, denaturation at 96°C was performed for 2 min, followed by 40 cycles of 96°C for 30 sec,

68°C for 15 sec and 72°C for 3 min, and finally by 72°C for 4 min. For Tb38-1 denaturation at 94°C for 2 min was followed by 10 cycles of 96°C for 15 sec, 68°C for 15 sec and 72°C for 1.5 min, 30 cycles of 96°C for 15 sec, 64°C for 15 sec and 72°C for 1.5, and finally by 72°C for 4 min.

The TbRa3 PCR fragment was digested with NdeI and EcoRI and cloned directly into pT7^L2 IL 1 vector using NdeI and EcoRI sites. The 38 kD PCR fragment was digested with Sse8387I, treated with T4 DNA polymerase to make blunt ends and then digested with EcoRI for direct cloning into the pT7^L2Ra3-1 vector which was digested with StuI and EcoRI. The 38-1 PCR fragment was digested with Eco47III and EcoRI and directly subcloned into pT7^L2Ra3/38kD-17 digested with the same enzymes. The whole fusion was then transferred to pET28b using NdeI and EcoRI sites. The fusion construct was confirmed by DNA sequencing.

The expression construct was transformed to BLR pLys S *E. coli* (Novagen, Madison, WI) and grown overnight in LB broth with kanamycin (30 µg/ml) and chloramphenicol (34 µg/ml). This culture (12 ml) was used to inoculate 500 ml 2XYT with the same antibiotics and the culture was induced with IPTG at an OD560 of 0.44 to a final concentration of 1.2 mM. Four hours post-induction, the bacteria were harvested and sonicated in 20 mM Tris (8.0), 100 mM NaCl, 0.1% DOC, 20 µg/ml Leupeptin, 20 mM PMSF followed by centrifugation at 26,000 X g. The resulting pellet was resuspended in 8 M urea, 20 mM Tris (8.0), 100 mM NaCl and bound to Pro-bond nickel resin (Invitrogen, Carlsbad, CA). The column was washed several times with the above buffer then eluted with an imidazole gradient (50 mM, 100 mM, 500 mM imidazole was added to 8 M urea, 20 mM Tris (8.0), 100 mM NaCl). The eluates containing the protein of interest were then dialzyed against 10 mM Tris (8.0).

The DNA and amino acid sequences for the resulting fusion protein (hereinafter referred to as TbRa3-38 kD-Tb38-1) are provided in SEQ ID NO: 147 and 148, respectively.

A fusion protein containing the two antigens TbH-9 and Tb38-1 (hereinafter referred to as TbH9-Tb38-1) without a hinge sequence, was prepared using a similar

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procedure to that described above. The DNA sequence for the TbH9-Tb38-1 fusion protein is provided in SEQ ID NO: 151.

A fusion protein containing TbRa3, the antigen 38kD, Tb38-1 and DPEP was prepared as follows.

Each of the DNA constructs TbRa3, 38 kD and Tb38-1 were modified by PCR and cloned into vectors essentially as described above, with the primers PDM-69 (SEQ ID NO:145 and PDM-83 (SEQ ID NO: 200) being used for amplification of the Tb38-1A fragment. Tb38-1A differs from Tb38-1 by a Dral site at the 3' end of the coding region that keeps the final amino acid intact while creating a blunt restriction site that is in frame. The TbRa3/38kD/Tb38-1A fusion was then transferred to pET28b using NdeI and EcoR1 sites.

DPEP DNA was used to perform PCR using the primers PDM-84 and PDM-85 (SEQ ID NO: 201 and 202, respectively) and 1 µl DNA at 50 ng/µl. Denaturation at 94 °C was performed for 2 min, followed by 10 cycles of 96 °C for 15 sec, 68 °C for 15 sec and 72 °C for 1.5 min; 30 cycles of 96 °C for 15 sec, 64 °C for 15 sec and 72 °C for 1.5 min; and finally by 72 °C for 4 min. The DPEP PCR fragment was digested with EcoRI and Eco72I and clones directly into the pET28Ra3/38kD/38-1A construct which was digested with DraI and EcoRI. The fusion construct was confirmed to be correct by DNA sequencing. Recombinant protein was prepared as described above. The DNA and amino acid sequences for the resulting fusion protein (hereinafter referred to as TbF-2) are provided in SEQ ID NO: 203 and 204, respectively.

EXAMPLE 8

USE OF M. TUBERCULOSIS FUSION PROTEINS FOR SERODIAGNOSIS OF TUBERCULOSIS

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The effectiveness of the fusion protein TbRa3-38 kD-Tb38-1, prepared as described above, in the serodiagnosis of tuberculosis infection was examined by ELISA.

The ELISA protocol was as described above in Example 6, with the fusion protein being coated at 200 ng/well. A panel of sera was chosen from a group of tuberculosis patients previously shown, either by ELISA or by western blot analysis, to react with each of

the three antigens individually or in combination. Such a panel enabled the dissection of the serological reactivity of the fusion protein to determine if all three epitopes functioned with the fusion protein. As shown in Table 5, all four sera that reacted with TbRa3 only were detectable with the fusion protein. Three sera that reacted only with Tb38-1 were also detectable, as were two sear that reacted with 38 kD alone. The remaining 15 sera were all positive with the fusion protein based on a cut-off in the assay of mean negatives +3 standard deviations. This data demonstrates the functional activity of all three epitopes in the fusion protein.

Table 5

<u>Reactivity of Tri-Peptide Fusion Protein with Sera from M. Tuberculosis Patients</u>

Serum ID	Status	1	A and/or Wes	Fusion	Fusion	
	1		y with Individ		recombinant	Recombinan
01B93I-40	 	38kd	Tb38-1	TbRa3	OD 450	Status
	TB	-	<u> </u>	+	0.413	+
01B93I-41	TB	<u> </u>	+	+	0.392	+
01B93I-29	TB	+	-	+	2.217	
01B93I-109	TB	+	±	+	0.522	+
01B93I-132	ТВ	+	+	+	0.937	+
5004	TB	±	+	±		+
15004	ТВ	+	+		1.098	+
39004	TB	+	+	+	2.077	+
68004	TB	+	f	+	1.675	+
99004	TB	 	+	+	2.388	+
107004			+	±	0.607	+
92004	TB	-	+	±	0.667	+
	TB	+	± '	±	1.070	+
97004	TB	+	-	. ±	1.152	+
118004	TB	+	_	±	2.694	+
173004	TB	+	+	+	3.258	
175004	TB	+	_	+	2.514	+
274004	TB	_		+		+
276004	TB	_	+		3.220	+
282004	TB	+			2.991	+
<u>-</u>					0.824	+

				, 		
289004	ТВ	-	-	+	0.848	+
308004	ТВ	-	+	-	3.338	+
314004	ТВ	•	+	-	1.362	+
317004	ТВ	+	-	-	0.763	+
312004	ТВ	•	-	+	1.079	+
D176	PPD	-	_	_	0.145	
D162	PPD	-	-	-	0.073	-
D161	PPD	-	-	-	0.097	-
D27	PPD	_	-	_	0.082	-
A6-124	NORMAL	-	-	-	0.053	-
A6-125	NORMAL	-	-	_	0.087 /	-
A6-126	NORMAL	-	-	-	0.346/	<u>±</u>
A6-127	NORMAL	-	-	-	0.064	, -
A6-128	NORMAL	-	-		0.034	-
A6-129	NORMAL	-	-	-	0.037	-
-A6-130	NORMAL		-		0.057	-
A6-131	NORMAL				0.054	-
A6-132	NORMAL	-	-		0.022	_
A6-133	NORMAL	-	-		0.147	-
A6-134	NORMAL	-	<u>-</u>	-	0.101	-
A6-135	NORMAL	-	-		0.066	-
A6-136	NORMAL	-	-		0.054	-
A6-137	NORMAL	-	-	-	0.065	_
A6-138	NORMAL	-	-	-	0.041	-
A6-139	NORMAL	-	-		0.103	-
A6-140	NORMAL	-	-	_	0.212	-
A6-141	NORMAL	_	-	-	0.056	-
A6-142	NORMAL	-	-		0.051	-

The reactivity of the fusion protein TbF-2 with sera from *M. tuberculosis*infected patients was examined by ELISA using the protocol described above. The results of
these studies (Table 6) demonstrate that all four antigens function independently in the fusion
protein.

TABLE 6

REACTIVITY OF TBF-2 FUSION PROTEIN WITH TB AND NORMAL SERA

Serum ID	Status	TbF OD450	Status	TbF-2 OD450	Status	ELISA Reactivity			
						38 kD	TbRa3	Tb38-1	DPEP
B931-40	ТВ	0.57	+	0.321	+	•	+	-	+
B931-41	ТВ	0.601	+	0.396	+	+	+	+	-
B931-109	TB	0.494	+	0.404	+	+	+	±	•
B931-132	ТВ	1.502	+	1.292	+	+	+	+	±
5004	ТВ	1.806	+	1.666	+	±	±	+	-
15004	TB	2.862	+	2.468	+	+	+	+	-
39004	TB	2.443	+	1.722	+	+	+	+	•
68004	ТВ	2.871	+	2.575	+	+	+	+	-
99004	TB	0.691	+	0.971	+	-	±	+ ;	-
107004	TB	0.875	+	0.732	+	-	±	+ .	-
92004	TB	1.632	+	1.394	+	+	±	±	-
97004	ТВ	1.491	+	1.979	+	+	±	•	+
118004	TB	3.182	+	3.045	+	+	±	-	-
173004	ТВ	3.644	+	3.578	+	+	+	+	-
175004	TB	3.332	+	2.916	+	+	+	-	-
274004	TB	3.696	+	3.716	+	<u> </u>	+	-	+
276004	TB	3.243	+	2.56	+		-	+	-
282004	TB	1.249	+	1.234	+	+	<u> </u>	-	-
289004	TB	1.373	+	1.17	+		+	-	-
308004	ТВ	3.708	+	3.355	+	•	-	+	-
314004	TB	1.663	+	1.399	+	-	-	+	-
317004	TB	1.163	+	0.92	+	+	-	-	-
312004	ТВ	1.709	+	1.453	+	-	+	<u> </u>	<u> </u>
380004	TB	0.238	-	0.461	+	-		-	+
451004	TB	0.18	-	0.2		-	-	-	±
478004	TB	0.188	-	0.469	+	-	-	<u> </u>	<u>+</u>
410004	TB	0.384	+	2.392	+	<u> </u> ±	-	ļ	+
411004	TB	0.306	+	0.874	+	<u> - </u>	+	<u> </u>	+
421004	TB	0.357	+	1.456	+	<u> - </u>	+	<u> - </u>	+
528004	TB	0.047	-	0.196	-	<u> </u>	<u> - </u>	<u> </u>	+
A6-87	Normal	0.094	-	0.063	<u> </u> -	-	ļ -	<u> </u>	-
A6-88	Normal	0.214		0.19	<u> -</u>	-	<u> </u>	<u> - </u>	<u> </u>
A6-89	Normal	0.248	-	0.125	-		<u> - </u>	<u> </u>	<u> </u>
A6-90	Normal	0.179	-	0.206	-	-	<u> - </u>	ļ -	<u> </u>
A6-91	Normal	0.135	-	0.151	-	-	-	<u> </u>	<u> </u>
A6-92	Normal	0.064	-	0.097	-		<u> - </u>	ļ	<u> -</u>
A6-93	Normal	0.072	-	0.098	-	-	-	<u> </u>	-
A6-94	Normal	0.072	-	0.064	-	-	 -	<u> </u>	-
A6-95	Normal	0.125	Ī	0.159	-	-	<u> </u>	<u> - </u>	-
A6-96	Normal	0.121	-	0.12	-	-	-	<u> -</u>	<u> </u>
						<u> </u>			<u> </u>
Cut-off		0.284		0.266				<u></u>	<u> </u>

One of skill in the art will appreciate that the order of the individual antigens within the fusion protein may be changed and that comparable activity would be expected provided each of the epitopes is still functionally available. In addition, truncated forms of the proteins containing active epitopes may be used in the construction of fusion proteins.

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From the foregoing, it will be appreciated that, although specific embodiments of the invention have been described herein for the purpose of illustration, various modifications may be made without deviating from the spirit and scope of the invention.

SEQUENCE LISTING

(1) GENERAL INFORMATION:

- (i) APPLICANTS: Reed, Steven G.
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- (ii) TITLE OF INVENTION: COMPOUNDS AND METHODS FOR DIAGNOSIS OF TUBERCULOSIS
- (iii) NUMBER OF SEQUENCES: 209
 - (iv) CORRESPONDENCE ADDRESS:
 - (A) ADDRESSEE: SEED and BERRY LLP
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 - (C) CITY: Seattle
 - (D) STATE: Washington
 - (E) COUNTRY: USA
 - (F) ZIP: 98104-7092
 - (v) COMPUTER READABLE FORM:
 - (A) MEDIUM TYPE: Floppy disk
 - (B) COMPUTER: IBM PC compatible
 - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
 - (D) SOFTWARE: PatentIn Release #1.0, Version #1.30
 - (vi) CURRENT APPLICATION DATA:
 - (A) APPLICATION NUMBER:
 - (B) FILING DATE: 01-OCT-1997
 - (C) CLASSIFICATION:
- (viii) ATTORNEY/AGENT INFORMATION:
 - (A) NAME: Maki, David J.
 - (B) REGISTRATION NUMBER: 31,392
 - (C) REFERENCE/DOCKET NUMBER: 210121.417C7
 - (ix) TELECOMMUNICATION INFORMATION:
 - (A) TELEPHONE: (206) 622-4900
 - (B) TELEFAX: (206) 682-6031
- (2) INFORMATION FOR SEQ ID NO:1:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 766 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

CGAGGCACCG	GTAGTTTGAA	CCAAACGCAC	AATCGACGGG	CAAACGAACG	GAAGAACACA	60
ACCATGAAGA	TGGTGAAATC	GATCGCCGCA	GGTCTGACCG	CCGCGGCTGC	AATCGGCGCC	120
GCTGCGGCCG	GTGTGACTTC	GATCATGGCT	GGCGGCCCGG	TCGTATACCA	GATGCAGCCG	180
GTCGTCTTCG	GCGCGCCACT	GCCGTTGGAC	CCGGCATCCG	CCCCTGACGT	CCCGACCGCC	240
GCCCAGTTGA	CCAGCCTGCT	CAACAGCCTC	GCCGATCCCA	ACGTGTCGTT	TGCGAACAAG	300
GGCAGTCTGG	TCGAGGGCGG	CATCGGGGGC	ACCGAGGCGC	GCATCGCCGA	CCACAAGCTG	360
AAGAAGGCCG	CCGAGCACGG	GGATCTGCCG	CTGTCGTTCA	GCGTGACGAA	CATCCAGCCG	420
ececcecce	GTTCGGCCAC	CGCCGACGTT	TCCGTCTCGG	GTCCGAAGCT	CTCGTCGCCG	480
GTCACGCAGA	ACGTCACGTT	CGTGAATCAA	GGCGGCTGGA	TGCTGTCACG	CGCATCGGCG	540
ATGGAGTTGC	TGCAGGCCGC	AGGGNAACTG	ATTGGCGGGC	CGGNTTCAGC	CCGCTGTTCA	600
GCTACGCCGC	CCGCCTGGTG	ACGCGTCCAT	GTCGAACACT	CGCGCGTGTA	GCACGGTGCG	660
GTNTGCGCAG	GGNCGCACGC	ACCGCCCGGT	GCAAGCCGTC	CTCGAGATAG	GTGGTGNCTC	720
GNCACCAGNG	ANCACCCCCN	NNTCGNCNNT	TCTCGNTGNT	GNATGA		766

(2) INFORMATION FOR SEQ ID NO:2:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 752 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

ATGCATCACC ATCACCATCA CGATGAAGTC ACGGTAGAGA CGACCTCCGT CTTCCGCGCA 60

GACTTCCTCA GCGAGCTGGA CGCTCCTGCG CAAGCGGGTA CGGAGAGCGC GGTCTCCGGG 120

GTGGAAGGGC TCCCGCCGGG CTCGGCGTTG CTGGTAGTCA AACGAGGCCC CAACGCCGGG 180

TCCCGGTTCC TACTCGACCA AGCCATCACG TCGGCTGGTC GGCATCCCGA CAGCGACATA 240

TTTCTCGACG	ACGTGACCGT	GAGCCGTCGC	CATGCTGAAT	TCCGGTTGGA	AAACAACGAA	300
TTCAATGTCG	TCGATGTCGG	GAGTCTCAAC	GGCACCTACG	TCAACCGCGA	GCCCGTGGAT	360
rcggcggtgc	TGGCGAACGG	CGACGAGGTC	CAGATCGGCA	AGCTCCGGTT	GGTGTTCTTG	420
ACCGGACCCA	AGCAAGGCGA	GGATGACGGG	AGTACCGGGG	GCCCGTGAGC	GCACCCGATA	480
GCCCGCGCT	GGCCGGGATG	TCGATCGGGG	CGGTCCTCCG	ACCTGCTACG	ACCGGATTTT	540
CCCTGATGTC	CACCATCTCC	AAGATTCGAT	TCTTGGGAGG	CTTGAGGGTC	NGGGTGACCC	600
cccccccccc	CTCATTCNGG	GGTNTCGGCN	GGTTTCACCC	CNTACCNACT	GCCNCCCGGN	660
TTGCNAATTC	NTTCTTCNCT	GCCCNNAAAG	GGACCNTTAN	CTTGCCGCTN	GAAANGGTNA	720
TCCNGGGCCC	NTCCTNGAAN	CCCCNTCCCC	CT		1	752

(2) INFORMATION FOR SEQ ID NO:3:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 813 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

CATATGCATC ACCATCACCA	TCACACTTCT	AACCGCCCAG	CGCGTCGGGG	GCGTCGAGCA	60
CCACGCGACA CCGGGCCCGA	TCGATCTGCT	AGCTTGAGTC	TGGTCAGGCA	TCGTCGTCAG	120
CAGCGCGATG CCCTATGTTT	GTCGTCGACT	CAGATATCGC	GGCAATCCAA	TCTCCCGCCT	180
GCGGCCGGCG GTGCTGCAAA	CTACTCCCGG	AGGAATTTCG	ACGTGCGCAT	CAAGATCTTC	240
ATGCTGGTCA CGGCTGTCGT	TTTGCTCTGT	TGTTCGGGTG	TGGCCACGGC	CGCGCCCAAG	300
ACCTACTGCG AGGAGTTGAA	AGGCACCGAT	ACCGGCCAGG	CGTGCCAGAT	TCAAATGTCC	360
GACCCGGCCT ACAACATCAA	CATCAGCCTG	CCCAGTTACT	ACCCCGACCA	GAAGTCGCTG	420
GAAAATTACA TCGCCCAGAC	GCGCGACAAG	TTCCTCAGCG	CGGCCACATC	GTCCACTCCA	480
CGCGAAGCCC CCTACGAATT	GAATATCACC	TCGGCCACAT	ACCAGTCCGC	GATACCGCCG	540
CGTGGTACGC AGGCCGTGGT	GCTCAMGGTC	TACCACAACG	CCGGCGGCAC	GCACCCAACG	600
ACCACGTACA AGGCCTTCGA	TTGGGACCAG	GCCTATCGCA	AGCCAATCAC	CTATGACACG	660
CTGTGGCAGG CTGACACCGA	TCCGCTGCCA	GTCGTCTTCC	CCATTGTTGC	AAGGTGAACT	720

GAGCAACGCA GACCGGGACA ACWGGTATCG ATAGCCGCCN	AATGCCGGCT	TGGAACCCNG	780
TGAAATTATC ACAACTTCGC AGTCACNAAA NAA			813
(2) INFORMATION FOR SEQ ID NO:4:			
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 447 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 		·	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:			
CGGTATGAAC ACGGCCGCGT CCGATAACTT CCAGCTGTCC	CAGGGTGGGC	AGGGATTCGC	, 60
CATTCCGATC GGGCAGGCGA TGGCGATCGC GGGCCAGATC	CGATCGGGTG	GGGGGTCACC	120
CACCGTTCAT ATCGGGCCTA CCGCCTTCCT CGGCTTGGGT	GTTGTCGACA	ACAACGGCAA	180
CGGCGCACGA GTCCAACGCG TGGTCGGGAG CGCTCCGGCG	GCAAGTCTCG	GCATCTCCAC	240
CGGCGACGTG ATCACCGCGG TCGACGGCGC TCCGATCAAC	TCGGCCACCG	CGATGGCGGA	300
CGCGCTTAAC GGGCATCATC CCGGTGACGT CATCTCGGTG	AACTGGCAAA	CCAAGTCGGG	360
CGGCACGCGT ACAGGGAACG TGACATTGGC CGAGGGACCC	CCGGCCTGAT	TTCGTCGYGG	420
ATACCACCCG CCGGCCGGCC AATTGGA			447
(2) INFORMATION FOR SEQ ID NO:5:			
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 604 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 			
		•	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:			
GTCCCACTGC GGTCGCCGAG TATGTCGCCC AGCAAATGTC	TGGCAGCCGC	CCAACGGAAT	60
CCGGTGATCC GACGTCGCAG GTTGTCGAAC CCGCCGCCGC	GGAAGTATCG	GTCCATGCCT	120
AGCCCGGCGA CGGCGAGCGC CGGAATGGCG CGAGTGAGGA	GGCGGGCAAT	TTGGCGGGGC	180

CCGGCGA	CGG	NGAGCGCCGG	AATGGCGCGA	GTGAGGAGGT	GGNCAGTCAT	GCCCAGNGTG	240
ATCCAAT	CAA	CCTGNATTCG	GNCTGNGGGN	CCATTTGACA	ATCGAGGTAG	TGAGCGCAAA	30Ó
TGAATGA	TGG	AAAACGGGNG	GNGACGTCCG	NTGTTCTGGT	GGTGNTAGGT	GNCTGNCTGG	360
NGTNGNG	GNT	ATCAGGATGT	TCTTCGNCGA	AANCTGATGN	CGAGGAACAG	GGTGTNCCCG	420
NNANNCCI	NAN	GGNGTCCNAN	CCCNNNNTCC	TCGNCGANAT	CANANAGNCG	NTTGATGNGA	480
NAAAAGGG	STG	GANCAGNNNN	AANTNGNGGN	CCNAANAANC	NNNANNGNNG	NNAGNTNGNT	540
NNNTNTTN	INC	ANNNNNNTG	NNGNNGNNCN	NNNCAANCNN	NTNNNNGNAA	NNGGNTTNTT	600
NAAT							
							604

(2) INFORMATION FOR SEQ ID NO:6:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 633 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

TTGCANGTCG	AACCACCTCA	CTAAAGGGAA	A CAAAAGCTNG	AGCTCCACCG	CGGTGGCGGC	60
CGCTCTAGAA	CTAGTGKATM	YYYCKGGCTG	CAGSAATYCG	GYACGAGCAT	TAGGACAGTC	120
TAACGGTCCT	GTTACGGTGA	TCGAATGACC	GACGACATCC	TGCTGATCGA	CACCGACGAA	180
CGGGTGCGAA	CCCTCACCCT	CAACCGGCCG	CAGTCCCGYA	ACGCGCTCTC	GGCGGCGCTA	240
CGGGATCGGT	TTTTCGCGGY	GTTGGYCGAC	GCCGAGGYCG	ACGACGACAT	CGACGTCGTC	300
ATCCTCACCG	GYGCCGATCC	GGTGTTCTGC	GCCGGACTGG	ACCTCAAGGT	AGCTGGCCGG	360
GCAGACCGCG	CTGCCGGACA	TCTCACCGCG	GTGGGCGGCC	ATGACCAAGC	CGGTGATCGG	420
CGCGATCAAC	GGCGCCGCGG	TCACCGGCGG	GCTCGAACTG	GCGCTGTACT	GCGACATCCT	480
GATCGCCTCC (540
CTGGGGACTC A						
CCTGACCGGC G					COLGGALGAG	600
			000			633

(2) INFORMATION FOR SEQ ID NO:7:

ID INO ADSCEASANT .

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1362 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:

CGACGACGAC	GGCGCCGGAG	AGCGGGCGCG	AACGGCGATC	GACGCGGCCC	TGGCCAGAGT	60
CGGCACCACC	CAGGAGGGAG	TCGAATCATG	AAATTTGTCA	ACCATATTGA	GCCCGTCGCG	120
CCCCGCCGAG	CCGGCGGCGC	GGTCGCCGAG	GTCTATGCCG	AGGCCCGCCG	CGAGTTCGGC	180
CGGCTGCCCG	AGCCGCTCGC	CATGCTGTCC	CCGGACGAGG	GACTGCTCAC	CGCCGGCTGG	240
GCGACGTTGC	GCGAGACACT	GCTGGTGGGC	CAGGTGCCGC	GTGGCCGCAA	GGAAGCCGTC	300
GCCGCCGCCG	TCGCGGCCAG	CCTGCGCTGC	CCCTGGTGCG	TCGACGCACA	CACCACCATG	360
CTGTACGCGG	CAGGCCAAAC	CGACACCGCC	GCGGCGATCT	TGGCCGGCAC	AGCACCTGCC	420
GCCGGTGACC	CGAACGCGCC	GTATGTGGCG	TGGGCGGCAG	GAACCGGGAC	ACCGGCGGGA	480
CCGCCGGCAC	CGTTCGGCCC	GGATGTCGCC	GCCGAATACC	TGGGCACCGC	GGTGCAATTC	540
CACTTCATCG	CACGCCTGGT	CCTGGTGCTG	CTGGACGAAA	CCTTCCTGCC	GGGGGCCCG	600
CGCGCCCAAC	AGCTCATGCG	CCGCGCCGGT	GGACTGGTGT	TCGCCCGCAA	GGTGCGCGCG	660
GAGCATCGGC	CGGGCCGCTC	CACCCGCCGG	CTCGAGCCGC	GAACGCTGCC	CGACGATCTG	720
GCATGGGCAA	CACCGTCCGA	GCCCATAGCA	ACCGCGTTCG	CCGCGCTCAG	CCACCACCTG	780
GACACCGCGC	CGCACCTGCC	GCCACCGACT	CGTCAGGTGG	TCAGGCGGGT	CGTGGGGTCG	840
TGGCACGGCG	AGCCAATGCC	GATGAGCAGT	CGCTGGACGA	ACGAGCACAC	CGCCGAGCTG	900
CCCGCCGACC	TGCACGCGCC	CACCCGTĆTT	GCCCTGCTGA	CCGGCCTGGC	CCCGCATCAG	960
GTGACCGACG	ACGACGTCGC	CGCGGCCCGA	TCCCTGCTCG	ACACCGATGC	GGCGCTGGTT	1020
GGCGCCCTGG	CCTGGGCCGC	CTTCACCGCC	GCGCGGCGCA	TCGGCACCTG	GATCGGCGCC	1080
GCCGCCGAGG	GCCAGGTGTC	GCGGCAAAAC	CCGACTGGGT	GAGTGTGCGC	GCCCTGTCGG	1140
TAGGGTGTCA	TCGCTGGCCC	GAGGGATCTC	GCGGCGGCGA	ACGGAGGTGG	CGACACAGGT	1200
GGAAGCTGCG	CCCACTGGCT	TGCGCCCCAA	CGCCGTCGTG	GGCGTTCGGT	TGGCCGCACT	1260
GGCCGATCAG	GTCGGCGCCG	GCCCTTGGCC	GAAGGTCCAG	CTCAACGTGC	CGTCACCGAA	1320

PCT/US97/18214

GGACCGGACG GTCACCGGGG GTCACCCTGC GCGCCCAAGG AA

1362

(2) INFORMATION FOR SEQ ID NO:8:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1458 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

GCGACGACCC	CGATATGCCG	GGCACCGTAG	CGAAAGCCGT	CGCCGACGCA	CTCGGGCGCG	60
GTATCGCTCC	CGTTGAGGAC	ATTCAGGACT	GCGTGGAGGC	CCGGCTGGGG	GAAGCCGGTC	120
TGGATGACGT	GGCCCGTGTT	TACATCATCT	ACCGGCAGCG	GCGCGCCGAG	CTGCGGACGG	180
CTAAGGCCTT	GCTCGGCGTG	CGGGACGAGT	TAAAGCTGAG	CTTGGCGGCC	GTGACGGTAC	240
TGCGCGAGCG	CTATCTGCTG	CACGACGAGC	AGGGCCGGCC	GGCCGAGTCG	ACCGGCGAGC	300
TGATGGACCG	ATCGGCGCGC	TGTGTCGCGG	CGGCCGAGGA	CCAGTATGAG	CCGGGCTCGT	360
CGAGGCGGTG	GGCCGAGCGG	TTCGCCACGC	TATTACGCAA	CCTGGAATTC	CTGCCGAATT	420
CGCCCACGTT	GATGAACTCT	GGCACCGACC	TGGGACTGCT	CGCCGGCTGT	TTTGTTCTGC	480
CGATTGAGGA	TTCGCTGCAA	TCGATCTTTG	CGACGCTGGG	ACAGGCCGCC	GAGCTGCAGC	540
GGGCTGGAGG	CGGCACCGGA	TATGCGTTCA	GCCACCTGCG	ACCCGCCGGG	GATCGGGTGG	600
CCTCCACGGG	CGGCACGGCC	AGCGGACCGG	TGTCGTTTCT	ACGGCTGTAT	GACAGTGCCG	660
CGGGTGTGGT	CTCCATGGGC	GGTCGCCGGC	GTGGCGCCTG	TATGGCTGTG	CTTGATGTGT	720
CGCACCCGGA	TATCTGTGAT	TTCGTCACCG	CCAAGGCCGA	ATCCCCCAGC	GAGCTCCCGC	780
ATTTCAACCT	ATCGGTTGGT	GTGACCGACG	CGTTCCTGCG	GGCCGTCGAA	CGCAACGGCC	840
TACACCGGCT	GGTCAATCCG	CGAACCGGCA	AGATCGTCGC	GCGGATGCCC	GCCGCCGAGC	900
TGTTCGACGC	CATCTGCAAA	GCCGCGCACG	CCGGTGGCGA	TCCCGGGCTG	GTGTTTCTCG	960
ACACGATCAA	TAGGGCAAAC	CCGGTGCCGG	GGAGAGGCCG	CATCGAGGCG	ACCAACCCGT	1020
GCGGGGAGGT	CCCACTGCTG	CCTTACGAGT	CATGTAATCT	CGGCTCGATC	AACCTCGCCC	1080
GGATGCTCGC	CGACGGTCGC	GTCGACTGGG	ACCGGCTCGA	GGAGGTCGCC	GGTGTGGCGG	1140
TGCGGTTCCT	TGATGACGTC	ATCGATGTCA	GCCGCTACCC	CTTCCCCGAA	CTGGGTGAGG	1200

CGGCCCGCGC CACCCGCAAG ATCGGGCTGG GAGTCATGGG TTTGGCGGAA CTGCTTGCCG 1260

CACTGGGTAT TCCGTACGAC AGTGAAGAAG CCGTGCGGTT AGCCACCCGG CTCATGCGTC 1320

GCATACAGCA GGCGGCGCAC ACGGCATCGC GGAGGCTGGC CGAAGAGCGG GGCGCATTCC 1380

CGGCGTTCAC CGATAGCCGG TTCGCGCGGT CGGGCCCGAG GCGCAACGCA CAGGTCACCT 1440

CCGTCGCTCC GACGGCCA 1458

(2) INFORMATION FOR SEQ ID NO:9:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 862 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:9:

ACGGTGTAAT	CGTGCTGGAT	CTGGAACCGC	GTGGCCCGCT	ACCTACCGAG	ATCTACTGGC	60
GGCGCAGGGG	GCTGGCCCTG	GGCATCGCGG	TCGTCGTAGT	CGGGATCGCG	GTGGCCATCG	120
TCATCGCCTT	CGTCGACAGC	AGCGCCGGTG	CCAAACCGGT	CAGCGCCGAC	AAGCCGGCCT	180
CCGCCCAGAG	CCATCCGGGC	TCGCCGGCAC	CCCAAGCACC	CCAGCCGGCC	GGGCAAACCG	240
AAGGTAACGC	CGCCGCGGCC	CCGCCGCAGG	GCCAAAACCC	CGAGACACCC	ACGCCCACCG	300
CCGCGGTGCA	GCCGCCGCCG	GTGCTCAAGG	AAGGGGACGA	TTGCCCCGAT	TCGACGCTGG	360
CCGTCAAAGG	TTTGACCAAC	GCGCCGCAGT	ACTACGTCGG	CGACCAGCCG	AAGTTCACCA	420
TGGTGGTCAC	CAACATCGGC	CTGGTGTCCT	GTAAACGCGA	CGTTGGGGCC	GCGGTGTTGG	480
CCGCCTACGT	TTACTCGCTG	GACAACAAGC	GGTTGTGGTC	CAACCTGGAC	TGCGCGCCCT	540
CGAATGAGAC	GCTGGTCAAG	ACGTTTTCCC	CCGGTGAGCA	GGTAACGACC	GCGGTGACCT	600
GGACCGGGAT	GGGATCGGCG	CCGCGCTGCC	CATTGCCGCG	GCCGGCGATC	GGGCCGGGCA	660
CCTACAATCT	CGTGGTACAA	CTGGGCAATC	TGCGCTCGCT	GCCGGTTCCG	TTCATCCTGA	720
ATCAGCCGCC	GCCGCCGCCC	GGGCCGGTAC	CCGCTCCGGG	TCCAGCGCAG	GCGCCTCCGC	780
CGGAGTCTCC	CGCGCAAGGC	GGATAATTAT	TGATCGCTGA	TGGTCGATTC	CGCCAGCTGT	840
GACAACCCCT	CGCCTCGTGC	CG				862

(2) INFORMATION FOR SEQ ID NO:10:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 622 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:10:

TTGATCAGCA	CCGGCAAGGC	GTCACATGCC	TCCCTGGGTG	TGCAGGTGAC	CAATGACAAA	60
GACACCCCGG	GCGCCAAGAT	CGTCGAAGTA	GTGGCCGGTG	GTGCTGCCGC	GAACGCTGGA	120
GTGCCGAAGG	GCGTCGTTGT	CACCAAGGTC	GACGACCGCC	CGATCAACAG	CGCGGACGCG	180
TTGGTTGCCG	CCGTGCGGTC	CAAAGCGCCG	GGCGCCACGG	TGGCGCTAAC	CTTTCAGGAT	240
CCCTCGGGCG	GTAGCCGCAC	AGTGCAAGTC	ACCCTCGGCA	AGGCGGAGCA	GTGATGAAGG	300
TCGCCGCGCA	GTGTTCAAAG	CTCGGATATA	CGGTGGCACC	CATGGAACAG	CGTGCGGAGT	360
TGGTGGTTGG	CCGGGCACTT	GTCGTCGTCG	TTGACGATCG	CACGGCGCAC	GGCGATGAAG	420
ACCACAGCGG	GCCGCTTGTC	ACCGAGCTGC	TCACCGAGGC	CGGGTTTGTT	GTCGACGGCG	480
TGGTGGCGGT	GTCGGCCGAC	GAGGTCGAGA	TCCGAAATGC	GCTGAACACA	GCGGTGATCG	540
GCGGGGTGGA	CCTGGTGGTG	TCGGTCGGCG	GGACCGGNGT	GACGNCTCGC	GATGTCACCC	600
CGGAAGCCAC	CCGNGACATT	CT				622

(2) INFORMATION FOR SEQ ID NO:11:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1200 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:11:

GGCGCAGCGG	TAAGCCTGTT	GGCCGCCGGC	ACACTGGTGT	TGACAGCATG	CGGCGGTGGC	60
ACCAACAGCT	CGTCGTCAGG	CGCAGGCGGA	ACGTCTGGGT	CGGTGCACTG	CGGCGGCAAG	120
AAGGAGCTCC	ACTCCAGCGG	CTCGACCGCA	CAAGAAAATG	CCATGGAGCA	GTTCGTCTAT	180

GCCTACGTGC	GATCGTGCCC	GGGCTACACG	TTGGACTACA	ACGCCAACGG	GTCCGGTGCC	240
GGGGTGACCC	AGTTTCTCAA	CAACGAAACC	GATTTCGCCG	GCTCGGATGT	CCCGTTGAAT	300
CCGTCGACCG	GTCAACCTGA	CCGGTCGGCG	GAGCGGTGCG	GTTCCCCGGC	ATGGGACCTG	360
CCGACGGTGT	TCGGCCCGAT	CGCGATCACC	TACAATATCA	AGGGCGTGAG	CACGCTGAAT	420
CTTGACGGAC	CCACTACCGC	CAAGATTTTC	AACGGCACCA	TCACCGTGTG	GAATGATCCA	480
CAGATCCAAG	CCCTCAACTC	CGGCACCGAC	CTGCCGCCAA	CACCGATTAG	CGTTATCTTC	540
CGCAGCGACA	AGTCCGGTAC	GTCGGACAAC	TTCCAGAAAT	ACCTCGACGG	TGTATCCAAC	600
GGGGCGTGGG	GCAAAGGCGC	CAGCGAAACG	TTCAGCGGGG	GCGTCGGCGT	CGGCGCCAGC	660
GGGAACAACG	GAACGTCGGC	CCTACTGCAG	ACGACCGACG	GGTCGATCAC	CTACAACGAG	720
TGGTCGTTTG	CGGTGGGTAA	GCAGTTGAAC	ATGGCCCAGA	TCATCACGTC	GGCGGGTCCG	780
GATCCAGTGG	CGATCACCAC	CGAGTCGGTC	GGTAAGACAA	TCGCCGGGGC	CAAGATCATG	840
GGACAAGGCA	ACGACCTGGT	ATTGGACACG	TCGTCGTTCT	ACAGACCCAC	CCAGCCTGGC	900
TCTTACCCGA	TCGTGCTGGC	GACCTATGAG	ATCGTCTGCT	CGAAATACCC	GGATGCGACG	960
ACCGGTACTG	CGGTAAGGGC	GTTTATGCAA	GCCGCGATTG	GTCCAGGCCA	AGAAGGCCTG	1020
GACCAATACG	GCTCCATTCC	GTTGCCCAAA	TCGTTCCAAG	CAAAATTGGC	GGCCGCGGTG	1080
AATGCTATTT	CTTGACCTAG	TGAAGGGAAT	TCGACGGTGA	GCGATGCCGT	TCCGCAGGTA	1140
GGGTCGCAAT	TTGGGCCGTA	TCAGCTATTG	CGGCTGCTGG	GCCGAGGCGG	GATGGGCGAG	1200

(2) INFORMATION FOR SEQ ID NO:12:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1155 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:12:

GCAAGCAGCT	GCAGGTCGTG	CTGTTCGACG	AACTGGGCAT	GCCGAAGACC	AAACGCACCA	60.
AGACCGGCTA	CACCACGGAT	GCCGACGCGC	TGCAGTCGTT	GTTCGACAAG	ACCGGGCATC	120
CGTTTCTGCA	ACATCTGCTC	GCCCACCGCG	ACGTCACCCG	GCTCAAGGTC	ACCGTCGACG	180

GTTGCTCCA	AGCGGTGGCC	GCCGACGGCC	GCATCCACAC	CACGTTCAAC	CAGACGATCG	240
CCGCGACCGG	CCGGCTCTCC	TCGACCGAAC	CCAACCTGCA	GAACATCCCG	ATCCGCACCG	300
ACGCGGGCCG	GCGGATCCGG	GACGCGTTCG	TGGTCGGGGA	CGGTTACGCC	GAGTTGATGA	360
CGGCCGACTA	CAGCCAGATC	GAGATGCGGA	TCATGGGGCA	CCTGTCCGGG	GACGAGGGCC	420
TCATCGAGGC	GTTCAACACC	GGGGAGGACC	TGTATTCGTT	CGTCGCGTCC	CGGGTGTTCG	480
GTGTGCCCAT	CGACGAGGTC	ACCGGCGAGT	TGCGGCGCCG	GGTCAAGGCG	ATGTCCTACG	540
GGCTGGTTTA	CGGGTTGAGC	GCCTACGGCC	TGTCGCAGCA	GTTGAAAATC	TCCACCGAGG	600
AAGCCAACGA	GCAGATGGAC	GCGTATTTCG	CCCGATTCGG	CGGGGTGCGC	GACTACCTGC	660
GCGCCGTAGT	CGAGCGGGCC	CGCAAGGACG	GCTACACCTC	GACGGTGCTG	GGCCGTCGCC	720
GCTACCTGCC	CGAGCTGGAC	AGCAGCAACC	GTCAAGTGCG	GGAGGCCGCC	GAGCGGGCGG	780
CGCTGAACGC	GCCGATCCAG	GGCAGCGCGG	CCGACATCAT	CAAGGTGGCC	ATGATCCAGG	840
TCGACAAGGC	GCTCAACGAG	GCACAGCTGG	CGTCGCGCAT	GCTGCTGCAG	GTCCACGACG	900
AGCTGCTGTT	CGAAATCGCC	CCCGGTGAAC	GCGAGCGGGT	CGAGGCCCTG	GTGCGCGACA	960
AGATGGGCGG	CGCTTACCCG	CTCGACGTCC	CGCTGGAGGT	GTCGGTGGGC	TACGGCCGCA	1020
GCTGGGACGC	GGCGGCGCAC	TGAGTGCCGA	GCGTGCATCT	GGGGCGGAA	TTCGGCGATT	1080
TTTCCGCCCT	GAGTTCACGC	TCGGCGCAAT	CGGGACCGAG	TTTGTCCAGC	GTGTACCCGT	1140
CGAGTAGCCT	CGTCA					1159

(2) INFORMATION FOR SEQ ID NO:13:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1771 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:13:

GAGCGCCGTC	TGGTGTTTGA	ACGGTTTTAC	CGGTCGGCAT	CGGCACGGGC	GTTGCCGGGT	60
TCGGGCCTCG	GGTTGGCGAT	CGTCAAACAG	GTGGTGCTCA	ACCACGGCGG	ATTGCTGCGC	120
ATCGAAGACA	CCGACCCAGG	CGGCCAGCCC	CCTGGAACGT	CGATTTACGT	GCTGCTCCCC	180

GGCCGTCGGA	TGCCGATTCC	GCAGCTTCCC	GGTGCGACGG	CTGGCGCTCG	GAGCACGGAC	240
ATCGAGAACT	CTCGGGGTTC	GGCGAACGTT	ATCTCAGTGG	AATCTCAGTC	CACGCGCGCA	300
ACCTAGTTGT	GCAGTTACTG	TTGAAAGCCA	CACCCATGCC	AGTCCACGCA	TGGCCAAGTT	360
GGCCCGAGTA	GTGGGCCTAG	TACAGGAAGA	GCAACCTAGC	GACATGACGA	ATCACCCACG	420
GTATTCGCCA	CCGCCGCAGC	AGCCGGGAAC	CCCAGGTTAT	GCTCAGGGGC	AGCAGCAAAC	480
GTACAGCCAG	CAGTTCGACT	GGCGTTACCC	ACCGTCCCCG	CCCCGCAGC	CAACCCAGTA	540
CCGTCAACCC	TACGAGGCGT	TGGGTGGTAC	CCGGCCGGGT	CTGATACCTG	GCGTGATTCC	600
GACCATGACG	CCCCTCCTG	GGATGGTTCG	CCAACGCCCT	CGTGCAGGCA	TGTTGGCCAT	660
CGGCGCGGTG	ACGATAGCGG	TGGTGTCCGC	CGGCATCGGC	GGCGCGGCCG	CATCCCTGGT	720
CGGGTTCAAC	CGGGCACCCG	CCGGCCCCAG	CGGCGGCCCA	GTGGCTGCCA	GCGCGGCGCC	780
AAGCATCCCC	GCAGCAAACA	TGCCGCCGGG.	GTCGGTCGAA	CAGGTGGCGG	CCAAGGTGGT	840
GCCCAGTGTC	GTCATGTTGG	AAACCGATCT	GGGCCGCCAG	TCGGAGGAGG	GCTCCGGCAT	900
CATTCTGTCT	GCCGAGGGGC	TGATCTTGAC	CAACAACCAC	GTGATCGCGG	CGGCCGCCAA	960
GCCTCCCCTG	GGCAGTCCGC	CGCCGAAAAC	GACGGTAACC	TTCTCTGACG	GGCGGACCGC	1020
ACCCTTCACG	GTGGTGGGGG	CTGACCCCAC	CAGTGATATC	GCCGTCGTCC	GTGTTCAGGG	1080
CGTCTCCGGG	CTCACCCCGA	TCTCCCTGGG	TTCCTCCTCG	GACCTGAGGG	TCGGTCAGCC	1140
GGTGCTGGCG	ATCGGGTCGC	CGCTCGGTTT	GGAGGGCACC	GTGACCACGG	GGATCGTCAG	1200
CGCTCTCAAC	CGTCCAGTGT	CGACGACCGG	CGAGGCCGGC	AACCAGAACA	A CCGTGCTGGA	1260
CGCCATTCAG	ACCGACGCCG	CGATCAACCC	CGGTAACTCC	GGGGGCGCGC	TGGTGAACAT	1320
GAACGCTCAA	CTCGTCGGAG	TCAACTCGGC	CATTGCCACG	CTGGGCGCG	G ACTCAGCCGA	1380
TGCGCAGAGC	GGCTCGATCG	GTCTCGGTTT	TGCGATTCCA	GTCGACCAG	G CCAAGCGCAT	1440
CGCCGACGAG	TTGATCAGCA	CCGGCAAGGC	GTCACATGCC	TCCCTGGGT	G TGCAGGTGAC	1500
CAATGACAAA	GACACCCCGG	GCGCCAAGAT	CGTCGAAGTA	GTGGCCGGT	G GTGCTGCCGC	1560
GAACGCTGGA	GTGCCGAAGG	GCGTCGTTGT	CACCAAGGTO	GACGACCGC	C CGATCAACAG	1620
CGCGGACGCG	TTGGTTGCCG	CCGTGCGGTC	CAAAGCGCCC	GGCGCCACG	G TGGCGCTAAC	1680
CTTTCAGGAT	CCCTCGGGCG	GTAGCCGCAC	AGTGCAAGT	ACCCTCGGC	A AGGCGGAGCA	1740
GTGATGAAGG	TCGCCGCGCA	GTGTTCAAAG	C			1771

(2) INFORMATION FOR SEQ ID NO:14:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1058 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:14:

CTCCACCGCG	GTGGCGGCCG	CTCTAGAACT	AGTGGATCCC	CCGGGCTGCA	GGAATTCGGC	60
ACGAGGATCC	GACGTCGCAG	GTTGTCGAAC	ccgccgccgc	GGAAGTATCG	GTCCATGCCT	120
AGCCCGGCGA	CGGCGAGCGC	CGGAATGGCG	CGAGTGAGGA	GGCGGGCAAT	TTGGCGGGC	180
CCGGCGACGG	CGAGCGCCGG	AATGGCGCGA	GTGAGGAGGC	GGGCAGTCAT	GCCCAGCGTG	240
ATCCAATCAA	CCTGCATTCG	GCCTGCGGGC	CCATTTGACA	ATCGAGGTAG	TGAGCGCAAA	300
TGAATGATGG	AAAACGGGCG	GTGACGTCCG	CTGTTCTGGT	GGTGCTAGGT	GCCTGCCTGG	360
CGTTGTGGCT	ATCAGGATGT	TCTTCGCCGA	AACCTGATGC	CGAGGAACAG	GGTGTTCCCG	420
TGAGCCCGAC	GGCGTCCGAC	CCCGCGCTCC	TCGCCGAGAT	CAGGCAGTCG	CTTGATGCGA	480
CAAAAGGGTT	GACCAGCGTG	CACGTAGCGG	TCCGAACAAC	CGGGAAAGTC	GACAGCTTGC	540
TGGGTATTAC	CAGTGCCGAT	GTCGACGTCC	GGGCCAATCC	GCTCGCGGCA	AAGGGCGTAT	600
GCACCTACAA	CGACGAGCAG	GGTGTCCCGT	TTCGGGTACA	AGGCGACAAC	ATCTCGGTGA	660
AACTGTTCGA	CGACTGGAGC	AATCTCGGCT	CGATTTCTGA	ACTGTCAACT	TCACGCGTGC	720
TCGATCCTGC	CGCTGGGGTG	ACGCAGCTGC	TGTCCGGTGT	CACGAACCTC	CAAGCGCAAG	780
GTACCGAAGT	GATAGACGGA	ATTTCGACCA	CCAAAATCAC	CGGGACCATC	CCCGCGAGCT	840
CTGTCAAGAT	GCTTGATCCT	GGCGCCAAGA	GTGCAAGGCC	GGCGACCGTG	TGGATTGCCC	900
AGGACGGCTC	GCACCACCTC	GTCCGAGCGA	GCATCGACCT	CGGATCCGGG	TCGATTCAGC	960
TCACGCAGTC	GAAATGGAAC	GAACCCGTCA	ACGTCGACTA	GGCCGAAGTT	GCGTCGACGC	1020
GTTGNTCGAA	ACGCCCTTGT	GAACGGTGTC	AACGGNAC			1058

(2) INFORMATION FOR SEQ ID NO:15:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 542 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single

71

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:15: GAATTCGGCA CGAGAGGTGA TCGACATCAT CGGGACCAGC CCCACATCCT GGGAACAGGC 60 GGCGGCGGAG GCGGTCCAGC GGGCGCGGGA TAGCGTCGAT GACATCCGCG TCGCTCGGGT 120 CATTGAGCAG GACATGGCCG TGGACAGCGC CGGCAAGATC ACCTACCGCA TCAAGCTCGA 180 . AGTGTCGTTC AAGATGAGGC CGGCGCAACC GCGCTAGCAC GGGCCGGCGA GCAAGACGCA 240 AAATCGCACG GTTTGCGGTT GATTCGTGCG ATTTTGTGTC TGCTCGCCGA GGCCTACCAG 300 GCGCGGCCCA GGTCCGCGTG CTGCCGTATC CAGGCGTGCA TCGCGATTCC GGCGGCCACG 360 CCGGAGTTAA TGCTTCGCGT CGACCCGAAC TGGGCGATCC GCCGGNGAGC TGATCGATGA 420 CCGTGGCCAG CCCGTCGATG CCCGAGTTGC CCGAGGAAAC GTGCTGCCAG GCCGGTAGGA 480 AGCGTCCGTA GGCGGCGGTG CTGACCGGCT CTGCCTGCGC CCTCAGTGCG GCCAGCGAGC 540 542 GG

(2) INFORMATION FOR SEQ ID NO:16:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 913 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:

CGGTGCCGCC C	GCGCCTCCG	TTGCCCCCAT	TGCCGCCGTC	GCCGATCAGC	TGCGCATCGC	60
CACCATCACC G	CCTTTGCCG	CCGGCACCGC	CGGTGGCGCC	GGGGCCGCCG	ATGCCACCGC	120
TTGACCCTGG C	CCGCCGGCGC	CGCCATTGCC	ATACAGCACC	CCGCCGGGGG	CACCGTTACC	180
GCCGTCGCCA C	CCGTCGCCGC	CGCTGCCGTT	TCAGGCCGGG	GAGGCCGAAT	GAACCGCCGC	240
CAAGCCCGCC G	GCCGGCACCG	TTGCCGCCTT	TTCCGCCCGC	CCCGCCGGCG	CCGCCAATTG	300
CCGAACAGCC A	AMGCACCGTT	GCCGCCAGCC	CCGCCGCCGT	TAACGGCGCT	GCCGGGCGCC	360
GCCGCCGGAC C	CCGCCATTAC	CGCCGTTCCC	GTTCGGTGCC	CCGCCGTTAC	CGGCGCCCC	420

GTTTGCCGCC	AATATTCGGC	GGGCACCGCC	AGACCCGCCG	GGGCCACCAT	TGCCGCCGGG	480
CACCGAAACA	ACAGCCCAAC	GGTGCCGCCG	GCCCGCCGT	TTGCCGCCAT	CACCGGCCAT	540
TCACCGCCAG	CACCGCCGTT	AATGTTTATG	AACCCGGTAC	CGCCAGCGCG	GCCCCTATTG	600
ccggcgccg	GAGNGCGTGC	ccgccgcgc	CGCCAACGCC	CAAAAGCCCG	GGGTTGCCAC	660
CGGCCCCGCC	GGACCCACCG	GTCCCGCCGA	TCCCCCGTT	GCCGCCGGTG	CCGCCGCCAT	720
TGGTGCTGCT	GAAGCCGTTA	GCGCCGGTTC	ĆGCSGGTTCC	GGCGGTGGCG	CCNTGGCCGC	780
cgccccccc	GTTGCCGTAC	AGCCACCCC	CGGTGGCGCC	GTTGCCGCCA	TTGCCGCCAT	840
TGCCGCCGTT	GCCGCCATTG	CCGCCGTTCC	CGCCGCCACC	GCCGGNTTGG	ccccccccc	900
CGCCGGCGGC	CGC					913

(2) INFORMATION FOR SEQ ID NO:17:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1872 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:17:

GACTACGTTG GTGTAGAAAA	ATCCTGCCGC	CCGGACCCTT	AAGGCTGGGA	CAATTTCTGA	60
TAGCTACCCC GACACAGGAG	GTTACGGGAT	GAGCAATTCG	CGCCGCCGCT	CACTCAGGTG	120
GTCATGGTTG CTGAGCGTGC	TGGCTGCCGT	CGGGCTGGGC	CTGGCCACGG	CGCCGGCCCA	180
GGCGGCCCCG CCGGCCTTGT	CGCAGGACCG	GTTCGCCGAC	TTCCCCGCGC	TGCCCCTCGA	240
CCCGTCCGCG ATGGTCGCCC	AAGTGGCGCC	ACAGGTGGTC	AACATCAACA	CCAAACTGGG	300
CTACAACAAC GCCGTGGGCG	CCGGGACCGG	CATCGTCATC	GATCCCAACG	GTGTCGTGCT	360
GACCAACAAC CACGTGATCG	CGGGCGCCAC	CGACATCAAT	GCGTTCAGCG	TCGGCTCCGG	420
CCAAACCTAC GGCGTCGATG	TGGTCGGGTA	TGACCGCACC	CAGGATGTCG	CGGTGCTGCA	480
GCTGCGCGGT GCCGGTGGCC	TGCCGTCGGC	GGCGATCGGT	GGCGGCGTCG	CGGTTGGTGA	540
GCCCGTCGTC GCGATGGGCA	ACAGCGGTGG	GCAGGGCGGA	ACGCCCCGTG	CGGTGCCTGG	600
CAGGGTGGTC GCGCTCGGCC	AAACCGTGCA	GGCGTCGGAT	TCGCTGACCG	GTGCCGAAGA	660

GACATTGAAC	GGGTTGATCC	AGTTCGATGC	CGCAATCCAG	CCCGGTGATT	CGGGCGGCC	720
CGTCGTCAAC	GGCCTAGGAC	AGGTGGTCGG	TATGAACACG	GCCGCGTCCG	ATAACTTCCA	780
GCTGTCCCAG	GGTGGGCAGG	GATTCGCCAT	TCCGATCGGG	CAGGCGATGG	CGATCGCGGG	840
CCAAATCCGA	TCGGGTGGGG	GGTCACCCAC	CGTTCATATC	GGGCCTACCG	CCTTCCTCGG	900
CTTGGGTGTT	GTCGACAACA	ACGGCAACGG	CGCACGAGTC	CAACGCGTGG	TCGGAAGCGC	960
TCCGGCGGCA	AGTCTCGGCA	TCTCCACCGG	CGACGTGATC	ACCGCGGTCG	ACGGCGCTCC	1020
GATCAACTCG	GCCACCGCGA	TGGCGGACGC	GCTTAACGGG	CATCATCCCG	GTGACGTCAT	1080
CTCGGTGAAC	TGGCAAACCA	AGTCGGGCGG	CACGCGTACA	GGGAACGTGA	CATTGGCCGA	1140
GGGACCCCCG	GCCTGATTTG	TCGCGGATAC	CACCCGCCGG	CCGGCCAATT	GGATTGGCGC	1200
CAGCCGTGAT	TGCCGCGTGA	GCCCCGAGT	TCCGTCTCCC	GTGCGCGTGG	CATTGTGGAA	1260
GCAATGAACG	AGGCAGAACA	CAGCGTTGAG	CACCCTCCCG	TGCAGGGCAG	TTACGTCGAA	1320
GGCGGTGTGG	TCGAGCATCC	GGATGCCAAG	GACTTCGGCA	GCGCCGCCGC	CCTGCCCGCC	1380
GATCCGACCT	GGTTTAAGCA	CGCCGTCTTC	TACGAGGTGC	TGGTCCGGGC	GTTCTTCGAC	1440
GCCAGCGCGG	ACGGTTCCGN	CGATCTGCGT	GGACTCATCG	ATCGCCTCGA	CTACCTGCAG	1500
TGGCTTGGCA	TCGACTGCAT	CTGTTGCCGC	CGTTCCTACG	ACTCACCGCT	GCGCGACGGC	1560
GGTTACGACA	TTCGCGACTT	CTACAAGGTG	CTGCCCGAAT	TCGGCACCGT	CGACGATTTC	1620
GTCGCCCTGG	TCGACACCGC	TCACCGGCGA	GGTATCCGCA	TCATCACCGA	CCTGGTGATG	1680
AATCACACCT	CGGAGTCGCA	CCCCTGGTTT	CAGGAGTCCC	GCCGCGACCC	AGACGGACCG	1740
TACGGTGACT	ATTACGTGTG	GAGCGACACC	AGCGAGCGCT	ACACCGACGC	CCGGATCATC	1800
TTCGTCGACA	CCGAAGAGTC	GAACTGGTCA	TTCGATCCTG	TCCGCCGACA	GTTNCTACTG	1860
GCACCGATTC	TT					187

(2) INFORMATION FOR SEQ ID NO:18:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1482 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

⁽xi) SEQUENCE DESCRIPTION: SEQ ID NO:18:

CTTCGCCGAA ACCTGATGCC GAGGAACAGG GTGTTCCCGT GAGCCCGACG GCGTCCGACC	60
CCGCGCTCCT CGCCGAGATC AGGCAGTCGC TTGATGCGAC AAAAGGGTTG ACCAGCGTGC	120
ACGTAGCGGT CCGAACAACC GGGAAAGTCG ACAGCTTGCT GGGTATTACC AGTGCCGATG	180
TCGACGTCCG GGCCAATCCG CTCGCGGCAA AGGGCGTATG CACCTACAAC GACGAGCAGG	240
GTGTCCCGTT TCGGGTACAA GGCGACAACA TCTCGGTGAA ACTGTTCGAC GACTGGAGCA	300
ATCTCGGCTC GATTTCTGAA CTGTCAACTT CACGCGTGCT CGATCCTGCC GCTGGGGTGA	360
CGCAGCTGCT GTCCGGTGTC ACGAACCTCC AAGCGCAAGG TACCGAAGTG ATAGACGGAA	420
TTTCGACCAC CAAAATCACC GGGACCATCC CCGCGAGCTC TGTCAAGATG CTTGATCCTG	480
GCGCCAAGAG TGCAAGGCCG GCGACCGTGT GGATTGCCCA GGACGGCTCG CACCACCTCG	540
TCCGAGCGAG CATCGACCTC GGATCCGGGT CGATTCAGCT CACGCAGTCG AAATGGAACG	600
AACCCGTCAA CGTCGACTAG GCCGAAGTTG CGTCGACGCG TTGCTCGAAA CGCCCTTGTG	660
AACGGTGTCA ACGGCACCCG AAAACTGACC CCCTGACGGC ATCTGAAAAT TGACCCCCTA	720
GACCGGGCGG TTGGTGGTTA TTCTTCGGTG GTTCCGGCTG GTGGGACGCG GCCGAGGTCG	780
CGGTCTTTGA GCCGGTAGCT GTCGCCTTTG AGGGCGACGA CTTCAGCATG GTGGACGAGG	840
CGGTCGATCA TGGCGGCAGC AACGACGTCG TCGCCGCCGA AAACCTCGCC CCACCGGCCG	900
AAGGCCTTAT TGGACGTGAC GATCAAGCTG GCCCGCTCAT ACCGGGAGGA CACCAGCTGG	960
AAGAAGAGGT TGGCGGCCTC GGGCTCAAAC GGAATGTAAC CGACTTCGTC AACCACCAGG	1020
AGCGGATAGC GGCCAAACCG GGTGAGTTCG GCGTAGATGC GCCCGGCGTG GTGAGCCTCG	1080
GCGAACCGTG CTACCCATTC GGCGGCGGTG GCGAACAGCA CCCGATGACC GGCCTGACAC	1140
GCGCGTATCG CCAGGCCGAC CGCAAGATGA GTCTTCCCGG TGCCAGGCGG GGCCCAAAAA	1200
CACGACGTTA TCGCGGGCGG TGATGAAATC CAGGGTGCCC AGATGTGCGA TGGTGTCGCG	1260
TTTGAGGCCA CGAGCATGCT CAAAGTCGAA CTCTTCCAAC GACTTCCGAA CCGGGAAGCG	1320
GGCGGCGCG ATGCGGCCCT CACCACCATG GGACTCCCGG GCTGACACTT CCCGCTGCAG	1380
GCAGGCGGCC AGGTATTCTT CGTGGCTCCA GTTCTCGGCG CGGGCGCGAT CGGCCAGCCG	1440
GGACACTGAC TCACGCAGGG TGGGAGCTTT CAATGCTCTT GT	1482

(2) INFORMATION FOR SEQ ID NO:19:

- ID AND HOTEEAEAD I -

⁽i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 876 base pairs

- (B) TYPE: nucleic acid(C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:19:

GAATTCGGCA	CGAGCCGGCG	ATAGCTTCTG	GCCGCGCCC	GACCAGATGG	CTCGAGGGTT	60
CGTGCTCGGG	GCCACCGCCG	GGCGCACCAC	CCTGACCGGT	GAGGGCCTGC	AACACGCCGA	120
CGGTCACTCG	TTGCTGCTGG	ACGCCACCAA	CCCGGCGGTG	GTTGCCTACG	ACCCGGCCTT	180
CGCCTACGAA	ATCGGCTACA	TCGNGGAAAG	CGGACTGGCC	AGGATGTGCG	GGGAGAACCC	240
GGAGAACATC	TTCTTCTACA	TCACCGTCTA	CAACGAGCCG	TACGTGCAGC	CGCCGGAGCC	300
GGAGAACTTC	GATCCCGAGG	GCGTGCTGGG	GGGTATCTAC	CGNTATCACG	CGGCCACCGA	360
GCAACGCACC	AACAAGGNGC	AGATCCTGGC	CTCCGGGGTA	GCGATGCCCG	CGGCGCTGCG	420
GGCAGCACAG	ATGCTGGCCG	CCGAGTGGGA	TGTCGCCGCC	GACGTGTGGT	CGGTGACCAG	480
TTGGGGCGAG	CTAAACCGCG	ACGGGGTGGT	CATCGAGACC	GAGAAGCTCC	GCCACCCGA	540
TCGGCCGGCG	GGCGTGCCCT	ACGTGACGAG	AGCGCTGGAG	AATGCTCGGG	GCCCGGTGAT	600
CGCGGTGTCG	GACTGGATGC	GCGCGGTCCC	CGAGCAGATC	CGACCGTGGG	TGCCGGGCAC	660
ATACCTCACG	TTGGGCACCG	ACGGGTTCGG	TTTTTCCGAC	ACTCGGCCCG	CCGGTCGTCG	720
TTACTTCAAC	ACCGACGCCG	AATCCCAGGT	TGGTCGCGGT	TTTGGGAGGG	GTTGGCCGGG	780
TCGACGGGTG	AATATCGACC	CATTCGGTGC	CGGTCGTGGG	CCGCCCGCCC	AGTTACCCGG	840
ATTCGACGAA	GGTGGGGGGT	TGCGCCCGAN	TAAGTT			876

(2) INFORMATION FOR SEQ ID NO:20:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1021 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:20:

ATCCCCCGG GCTGCAGGAA TTCGGCACGA GAGACAAAAT TCCACGCGTT AATGCAGGAA

					TTATTTCGAC	120
AGCGAAGAC	C TGCCGCAGT	T GGCGAAGCA	TTTTACAGCO	AAGCGGTCG	GGAACGAAAC	180
CATGCAATG	A TGCTCGTGC	A ACACCTGCT	C GACCGCGACC	TTCGTGTCG	AATTCCCGGC	240
GTAGACACG	G TGCGAAACC	A GTTCGACAGA	A CCCCGCGAGG	CACTGGCGCT	GGCGCTCGAT	300
CAGGAACGC	A CAGTCACCGA	CCAGGTCGG1	CGGCTGACAG	CGGTGGCCCG	CGACGAGGGC	360
GATTTCCTC	G GCGAGCAGTT	CATGCAGTGG	TTCTTGCAGG	AACAGATCGA	AGAGGTGGCC	420
TTGATGGCAA	A CCCTGGTGCG	GGTTGCCGAT	CGGGCCGGGG	CCAACCTGTT	CGAGCTAGAG	480
AACTTCGTCG	CACGTGAAGT	GGATGTGGCG	CCGGCCGCAT	CAGGCGCCCC	GCACGCTGCC	540
GGGGGCCGCC	TCTAGATCCC	TGGGGGGGAT	CAGCGAGTGG	TCCCGTTCGC	CCGCCCGTCT	600
TCCAGCCAGG	CCTTGGTGCG	GCCGGGGTGG	TGAGTACCAA	TCCAGGCCAC	CCCGACCTCC	660
CGGNAAAAGT	CGATGTCCTC	GTACTCATCG	ACGTTCCAGG	AGTACACCGC	CCGGCCCTGA	720
GCTGCCGAGC	GGTCAACGAG	TTGCGGATAT	TCCTTTAACG	CAGGCAGTGA	GGGTCCCACG	780
GCGGTTGGCC	CGACCGCCGT	GGCCGCACTG	CTGGTCAGGT	ATCGGGGGGT	CTTGGCGAGC	840
AACAACGTCG	GCAGGAGGGG	TGGAGCCCGC	CGGATCCGCA	GACCGGGGGG	GCGAAAACGA	900
CATCAACACC	GCACGGGATC	GATCTGCGGA	GGGGGGTGCG	GGAATACCGA	ACCGGTGTAG	960
GAGCGCCAGC	AGTTGTTTTT	CCACCAGCGA	AGCGTTTTCG	GGTCATCGGN	GGCNNTTAAG	1020
T						1021

(2) INFORMATION FOR SEQ ID NO:21:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 321 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:21:

CGTGCCGACG AACGGAAGAA CACAACCATG AAGATGGTGA AATCGATCGC CGCAGGTCTG 60
ACCGCCGCGG CTGCAATCGG CGCCGCTGCG GCCGGTGTGA CTTCGATCAT GGCTGGCGGN 120
CCGGTCGTAT ACCAGATGCA GCCGGTCGTC TTCGGCGCGC CACTGCCGTT GGACCCGGNA 180

77

TCCGCCCCTG	ANGTCCCGAC	CGCCGCCCAG	TGGACCAGNC	TGCTCAACAG	NCTCGNCGAT	240
CCCAACGTGT	CGTTTGNGAA	CAAGGGNAGT	CTGGTCGAGG	GNGGNATCGG	NGGNANCGAG	300
GGNGNGNATC	GNCGANCACA	A				321

(2) INFORMATION FOR SEQ ID NO:22:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 373 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:22:

TCTTATCGGT TCCGGTTGGC GACGGGTTTT GGGNGCGGGT GGTTAACCCG CTCGGCCAGC 60

CGATCGACGG GCGCGAGAC GTCGACTCCG ATACTCGGCG CGCGCTGGAG CTCCAGGCGC 120

CCTCGGTGGT GNACCGGCAA GGCGTGAAGG AGCCGTTGNA GACCGGGATC AAGGCGATTG 180

ACGCGATGAC CCCGATCGGC CGCGGGCAGC GCCAGCTGAT CATCGGGGAC CGCAAGACCG 240

GCAAAAACCG CCGTCTGTGT CGGACACCAT CCTCAAACCA GCGGGAAGAA CTGGGAGTCC 300

GGTGGATCCC AAGAAGCAGG TGCGCTTGTG TATACGTTGG CCATCGGGCA AGAAGGGGAA 360

CTTACCATCG CCG

(2) INFORMATION FOR SEQ ID NO:23:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 352 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:23:

GTGACGCCGT GATGGGATTC CTGGGCGGGG CCGGTCCGCT GGCGGTGGTG GATCAGCAAC 60

TGGTTACCCG GGTGCCGCAA GGCTGGTCGT TTGCTCAGGC AGCCGCTGTG CCGGTGGTGT 120

TCTTGACGGC CTGGTACGGG TTGGCCGATT TAGCCGAGAT CAAGGCGGGC GAATCGGTGC 180

TGATCCATGC CGGTACCGGC GGTGTGGCCA TGGCGGCTGT GCAGCTGGCT CGCCAGTGGG 240

GCGTGGAGGT	TTTCGTCACC	GCCAGCCGTG	GNAAGTGGGA	CACGCTGCGC	GCCATNGNGT	300
TTGACGACGA	NCCATATCGG	NGATTCCCNC	ACATNCGAAG	TTCCGANGGA	GA	352

(2) INFORMATION FOR SEQ ID NO:24:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 726 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:24: GAAATCCGCG TTCATTCCGT TCGACCAGCG GCTGGCGATA ATCGACGAAG TGATCAAGCC 60 GCGGTTCGCG GCGCTCATGG GTCACAGCGA GTAATCAGCA AGTTCTCTGG TATATCGCAC 120 CTAGCGTCCA GTTGCTTGCC AGATCGCTTT CGTACCGTCA TCGCATGTAC CGGTTCGCGT 180 GCCGCACGCT CATGCTGGCG GCGTGCATCC TGGCCACGGG TGTGGCGGGT CTCGGGGTCG 240 GCGCGCAGTC CGCAGCCCAA ACCGCGCCGG TGCCCGACTA CTACTGGTGC CCGGGGCAGC 300 CTTTCGACCC CGCATGGGG CCCAACTGGG ATCCCTACAC CTGCCATGAC GACTTCCACC 360 GCGACAGCGA CGGCCCCGAC CACAGCCGCG ACTACCCCGG ACCCATCCTC GAAGGTCCCG 420 TGCTTGACGA TCCCGGTGCT GCGCCGCCGC CCCCGGCTGC CGGTGGCGGC GCATAGCGCT 480 CGTTGACCGG GCCGCATCAG CGAATACGCG TATAAACCCG GGCGTGCCCC CGGCAAGCTA 540 CGACCCCGG CGGGGCAGAT TTACGCTCCC GTGCCGATGG ATCGCGCCGT CCGATGACAG 600 AAAATAGGCG ACGGTTTTGG CAACCGCTTG GAGGACGCTT GAAGGGAACC TGTCATGAAC 660 GGCGACAGCG CCTCCACCAT CGACATCGAC AAGGTTGTTA CCCGCACACC CGTTCGCCGG 720 726 ATCGTG

(2) INFORMATION FOR SEQ ID NO:25:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 580 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE	DESCRIPTION:	SEQ	ID	NO:25:
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CGCGACGACG	ACGAACGTCG	GGCCCACCAC	CGCCTATGCG	TTGATGCAGG	CGACCGGGAT	60
GGTCGCCGAC	CATATCCAAG	CATGCTGGGT	GCCCACTGAG	CGACCTTTTG	ACCAGCCGGG	120
CTGCCCGATG	GCGGCCCGGT	GAAGTCATTG	CGCCGGGGCT	TGTGCACCTG	ATGAACCCGA	180
ATAGGGAACA	ATAGGGGGGT	GATTTGGCAG	TTCAATGTCG	GGTATGGCTG	GAAATCCAAT	240
GGCGGGGCAT	GCTCGGCGCC	GACCAGGCTC	GCGCAGGCGG	GCCAGCCCGA	ATCTGGAGGG	300
AGCACTCAAT	GGCGGCGATG	AAGCCCCGGA	CCGGCGACGG	TCCTTTGGAA	GCAACTAAGG	360
AGGGGCGCGG	CATTGTGATG	CGAGTACCAC	TTGAGGGTGG	CGGTCGCCTG	GTCGTCGAGC	420
TGACACCCGA	CGAAGCCGCC	GCACTGGGTG	ACGAACTCAA	AGGCGTTACT	AGCTAAGACC	480
AGCCCAACGG	CGAATGGTCG	GCGTTACGCG	CACACCTTCC	GGTAGATGTC	CAGTGTCTGC	540
TCGGCGATGT	ATGCCCAGGA	GAACTCTTGG	ATACAGCGCT			580

(2) INFORMATION FOR SEQ ID NO:26:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 160 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:26:

AACGGAGGCG CCGGGGGTTT TGGCGGGGCC GGGGCGGTCG GCGGCAACGG CGGGGCCGGC 60

GGTACCGCCG GGTTGTTCGG TGTCGGCGGG GCCGGTGGGG CCGGAGGCAA CGGCATCGCC 120

GGTGTCACGG GTACGTCGGC CAGCACACCG GGTGGATCCG 160

(2) INFORMATION FOR SEQ ID NO:27:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 272 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:27:	
GACACCGATA CGATGGTGAT GTACGCCAAC GTTGTCGACA CGCTCGAGGC GTTCACGATC	60
CAGCGCACAC CCGACGGCGT GACCATCGGC GATGCGGCCC CGTTCGCGGA GGCGGCTGCC	120
AAGGCGATGG GAATCGACAA GCTGCGGGTA ATTCATACCG GAATGGACCC CGTCGTCGCT	180
GAACGCGAAC AGTGGGACGA CGGCAACAAC ACGTTGGCGT TGGCGCCCGG TGTCGTTGTC	240
GCCTACGAGC GCAACGTACA GACCAACGCC CG	272
(2) INFORMATION FOR SEQ ID NO:28:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 317 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:28: GCAGCCGGTG GTTCTCGGAC TATCTGCGCA CGGTGACGCA GCGCGACGTG CGCGAGCTGA	60
AGCGGATCGA GCAGACGGAT CGCCTGCCGC GGTTCATGCG CTACCTGGCC GCTATCACCG	60
CGCAGGAGCT GAACGTGGCC GAAGCGGCGC GGGTCATCGG GGTCGACGCG GGGACGATCC	120 180
GTTCGGATCT GGCGTGGTTC GAGACGGTCT ATCTGGTACA TCGCCTGCCC GCCTGGTCGC	240
GGAATCTGAC CGCGAAGATC AAGAAGCGGT CAAAGATCCA CGTCGTCGAC AGTGGCTTCG	300
CGGCCTGGTT GCGCGGG	317
(2) INFORMATION FOR SEQ ID NO:29:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 182 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:29:

GATCGTGGAG CTGTCGATGA ACAGCGTTGC CGGACGCGCG GCGGCCAGCA CGTCGGTGTA 60

GCAGCGCCGG ACCACCTCGC CGGTGGGCAG CATGGTGATG ACCACGTCGG CCTCGGCC	CAC 120					
CGCTTCGGGC GCGCTACGAA ACACCGCGAC ACCGTGCGCG GCGGCGCCGG ACGCCGCC	CGT 180					
GG	182					
(2) INFORMATION FOR SEQ ID NO:30:						
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 308 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 						
	ļ					
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:30:						
GATCGCGAAG TTTGGTGAGC AGGTGGTCGA CGCGAAAGTC TGGGCGCCTG CGAAGCG	GGT 60					
CGGCGTTCAC GAGGCGAAGA CACGCCTGTC CGAGCTGCTG CGGCTCGTCT ACGGCGG						
GAGGTTGAGA TTGCCCGCCG CGGCGAGCCG GTAGCAAAGC TTGTGCCGCT GCATCCT						
GAGACTCGGC GGTTAGGCAT TGACCATGGC GTGTACCGCG TGCCCGACGA TTTGGAC	GCT 240					
CCGTTGTCAG ACGACGTGCT CGAACGCTTT CACCGGTGAA GCGCTACCTC ATCGACA	ACCC 300					
ACGTTTGG	308					
(2) INFORMATION FOR SEQ ID NO:31: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 267 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear						
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:31:						
CCGACGACGA GCAACTCACG TGGATGATGG TCGGCAGCGG CATTGAGGAC GGAGAGA	AATC 60					
CGGCCGAAGC TGCCGCGCG CAAGTGCTCA TAGTGACCGG CCGTAGAGGG CTCCCCC	CGAT 120					
GGCACCGGAC TATTCTGGTG TGCCGCTGGC CGGTAAGAGC GGGTAAAAGA ATGTGAC	GGGG 180					
ACACGATGAG CAATCACACC TACCGAGTGA TCGAGATCGT CGGGACCTCG CCCGACG	GGCG 240					
TCGACGCGGC AATCCAGGGC GGTCTGG	267					

(2) INFORMATION FOR SEQ ID NO:32:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1539 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:32:

CTCGTGCCGA	AAGAATGTGA	GGGGACACGA	TGAGCAATCA	CACCTACCGA	GTGATCGAGA	60
TCGTCGGGAC	CTCGCCCGAC	GGCGTCGACG	CGGCAATCCA	GGGCGGTCTG	GCCCGAGCTG	120
CGCAGACCAT	GCGCGCGCTG	GACTGGTTCG	AAGTACAGTC	AATTCGAGGC	CACCTGGTCG	180
ACGGAGCGGT	CGCGCACTTC	CAGGTGACTA	TGAAAGTCGG	CTTCCGCTGG	AGGATTCCTG	240
AACCTTCAAG	CGCGGCCGAT	AACTGAGGTG	CATCATTAAG	CGACTTTTCC	AGAACATCCT	300
GACGCGCTCG	AAACGCGGTT	CAGCCGACGG	TGGCTCCGCC	GAGGCGCTGC	CTCCAAAATC	360
CCTGCGACAA	TTCGTCGGCG	GCGCCTACAA	GGAAGTCGGT	GCTGAATTCG	TCGGGTATCT	420
GGTCGACCTG	TGTGGGCTGC	AGCCGGACGA	AGCGGTGCTC	GACGTCGGCT	GCGGCTCGGG	480
GCGGATGGCG	TTGCCGCTCA	CCGGCTATCT	GAACAGCGAG	GGACGCTACG	CCGGCTTCGA	540
TATCTCGCAG	AAAGCCATCG	CGTGGTGCCA	GGAGCACATC	ACCTCGGCGC	ACCCCAACTT	600
CCAGTTCGAG	GTCTCCGACA	TCTACAACTC	GCTGTACAAC	CCGAAAGGGA	AATACCAGTC	660
ACTAGACTTT	CGCTTTCCAT	ATCCGGATGC	GTCGTTCGAT	GTGGTGTTTC	TTACCTCGGT	720
GTTCACCCAC	ATGTTTCCGC	CGGACGTGGA	GCACTATCTG	GACGAGATCT	CCCGCGTGCT	780
GAAGCCCGGC	GGACGATGCC	TGTGCACGTA	CTTCTTGCTC	AATGACGAGT	CGTTAGCCCA	840
CATCGCGGAA	GGAAAGAGTG	CGCACAACTT	CCAGCATGAG	GGACCGGGTT	ATCGGACAAT	900
CCACAAGAAG	CGGCCCGAAG	AAGCAATCGG	CTTGCCGGAG	ACCTTCGTCA	GGGATGTCTA	960
TGGCAAGTTC	GGCCTCGCCG	TGCACGAACC	ATTGCACTAC	GGCTCATGGA	GTGGCCGGGA	1020
ACCACGCCTA	AGCTTCCAGG	ACATCGTCAT	CGCGACCAAA	ACCGCGAGCT	AGGTCGGCAT	1080
CCGGGAAGCA	TCGCGACACC	GTGGCGCCGA	GCGCCGCTGC	CGGCAGGCCG	ATTAGGCGGG	1140
CAGATTAGCC	CGCCGCGGCT	CCCGGCTCCG	AGTACGGCGC	CCCGAATGGC	GTCACCGGCT	1200
GGTAACCACG	CTTGCGCGCC	TGGGCGGCGG	CCTGCCGGAT	CAGGTGGTAG	ATGCCGACAA	1260

(2) INFORMATION FOR SEQ ID NO:33:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 851 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:33:

CTGCAGGGTG	GCGTGGATGA	GCGTCACCGC	GGGGCAGGCC	GAGCTGACCG	CCGCCCAGGT	60
			GTATGGGCTG			120
			GATAGCGACC	•		180
						240
			CGGCGAGATG			
GATGTTTGGC	TACGCCGCGG	CGACGGCGAC	GGCGACGGCG	ACGTTGCTGC	CGTTCGAGGA	300
GGCGCCGGAG	ATGACCAGCG	CGGGTGGGCT	CCTCGAGCAG	GCCGCCGCGG	TCGAGGAGGC	360
CTCCGACACC	GCCGCGGCGA	ACCAGTTGAT	GAACAATGTG	CCCCAGGCGC	TGAAACAGTT	420
GGCCCAGCCC	ACGCAGGGCA	CCACGCCTTC	TTCCAAGCTG	GGTGGCCTGT	GGAAGACGGT	480
CTCGCCGCAT	CGGTCGCCGA	TCAGCAACAT	GGTGTCGATG	GCCAACAACC	ACATGTCGAT	540
GACCAACTCG	GGTGTGTCGA	TGACCAACAC	CTTGAGCTCG	ATGTTGAAGG	GCTTTGCTCC	600
GGCGGCGGCC	GCCCAGGCCG	TGCAAACCGC	GGCGCAAAAC	GGGGTCCGGG	CGATGAGCTC	660
GCTGGGCAGC	TCGCTGGGTT	CTTCGGGTCT	GGGCGGTGGG	GTGGCCGCCA	ACTTGGGTCG	720
GGCGGCCTCG	GTACGGTATG	GTCACCGGGA	TGGCGGAAAA	TATGCANAGT	CTGGTCGGCG	780
GAACGGTGGT	CCGGCGTAAG	GTTTACCCCC	GTTTTCTGGA	TGCGGTGAAC	TTCGTCAACG	840
GAAACAGTTA	С					851

(2) INFORMATION FOR SEQ ID NO:34:

131	SECUENCE	CHARACTERISTICS
	STANCE	CHAIGCIDITIOTICS

(A) LENGTH: 254 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:34:

GATCGATCGG	GCGGAAATTT	GGACCAGATT	CGCCTCCGGC	GATAACCCAA	TCAATCGAAC	60
CTAGATTTAT	TCCGTCCAGG	GGCCCGAGTA	ATGGCTCGCA	GGAGAGGAAC	CTTACTGCTG	120
CGGGCACCTG	TCGTAGGTCC	TCGATACGGC	GGAAGGCGTC	GACATTTTCC	ACCGACACCC	180
CCATCCAAAC	GTTCGAGGGC	CACTCCAGCT	TGTGAGCGAG	GCGACGCAGT	CGCAGGCTGC	240
GCTTGGTCAA	GATC					254

(2) INFORMATION FOR SEQ ID NO:35:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1227 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:35:

GATCCTGACC	GAAGCGGCCG	CCGCCAAGGC	GAAGTCGCTG	TTGGACCAGG	AGGGACGGGA	60
CGATCTGGCG	CTGCGGATCG	CGGTTCAGCC	GGGGGGTGC	GCTGGATTGC	GCTATAACCT	120
TTTCTTCGAC	GACCGGACGC	TGGATGGTGA	CCAAACCGCG	GAGTTCGGTG	GTGTCAGGTT	180
GATCGTGGAC	CGGATGAGCG	CGCCGTATGT	GGAAGGCGCG	TCGATCGATT	TCGTCGACAC	240
TATTGAGAAG	CAAGGTTCAC	CATCGACAAT	CCCAACGCCA	CCGGCTCCTG	CGCGTGCGGG	300
GATTCGTTCA	ACTGATAAAA	CGCTAGTACG	ACCCCGCGGT	GCGCAACACG	TACGAGCACA	360
CCAAGACCTG	ACCGCGCTGG	AAAAGCAACT	GAGCGATGCC.	TTGCACCTGA	CCGCGTGGCG	420
GGCCGCCGGC	GGCAGGTGTC	ACCTGCATGG	TGAACAGCAC	CTGGGCCTGA	TATTGCGACC	480
AGTACACGAT	TTTGTCGATC	GAGGTCACTT	CGACCTGGGA	GAACTGCTTG	CGGAACGCGT	540

CTTGGCCAAG	GCCTGATCGG	AGCGCTTGTC	GCGCACGCCG	TCGTGGATAC	600
ATTGCGAACG	ATGGTGTCCA	CATCGCGGTT	CTCCAGCGCG	TTGAGGTATC	660
GGTTTTGGCC	GGTCCCTCCG	AGAATGTGCC	TGCCGTGTTG	GCTCCGTTGG	720
GTATATGATC	GCCGCCGTCA	TAGCCGACAC	CAGCGCGAGG	GCTACCACAA	780
CAGCCGCTTG	TGCCGTCGCT	TCGGGTAGGA	CACCTGCGGC	GGCACGCCGG	840
GGGCGGCAGC	GCCGCGTCGT	CTGCCGGTCC	CGGGGCGAAG	GCCGGTTCGG	900
GTCGTGGGGG	TAGTCCAGGG	CTTGGGGTTC	GTGGGATGAG	GGCTCGGGGT	960
TCCGTTGGTG	CCGACACCGG	GGTTCGGCGA	GTGGGGACCG	GGCATTGTGG	1020
GTGGTGGACG	GGACCAGCTG	CTAGGGCGAC	AACCGCCCGT	CGCGTCAGCC	1080
GCAATCAGGT	GAGCTCCCTA	GGCAGGCTAG	CGCAACAGCT	GCCGTCAGCT	1140
CGGGGCGGC	CGCGGCGCCG	ATAATGTTGA	AAGACTAGGC	AACCTTAGGA	1200
GAGATTTTGT	GACGATC				1227
	ATTGCGAACG GGTTTTGGCC GTATATGATC CAGCCGCTTG GGGCGGCAGC GTCGTGGGGG TCCGTTGGTG GTGGTGGACG GTGGTGGACG GCAATCAGGT CGGGGGGGC	ATTGCGAACG ATGGTGTCCA GGTTTTGGCC GGTCCCTCG GTATATGATC GCCGCCGTCA CAGCCGCTTG TGCCGTCGCT GGGCGGCAGC GCCGCTCGT GTCGTGGGGG TAGTCCAGGG TCCGTTGGTG CCGACACCGG GTGGTGGACG GGACCAGCTG GCAATCAGGT GAGCTCCTA	ATTGCGAACG ATGGTGTCCA CATCGCGGTT GGTTTTGGCC GGTCCCTCCG AGAATGTGCC GTATATGATC GCCGCCGTCA TAGCCGACAC CAGCCGCTTG TGCCGTCGCT TCGGGTAGGA GGGCGGCAGC GCCGCTCGT CTGCCGTCC GTCGTGGGGG TAGTCCAGGG CTTGGGGTTC TCCGTTGGTG CCGACACCGG GGTTCGGCGA GTGGTGGACG GGACCAGCTG CTAGGGCGAC GCAATCAGGT GAGCTCCCTA GGCAGGCTAG CGGGGGGGGC CGCGGCGCC ATAATGTTGA	ATTGCGAACG ATGGTGTCCA CATCGCGGTT CTCCAGCGCG GGTTTTGGCC GGTCCCTCCG AGAATGTGCC TGCCGTGTTG GTATATGATC GCCGCCGTCA TAGCCGACAC CAGCGCGAGG CAGCCGCTTG TGCCGTCGCT TCGGGTAGGA CACCTGCGGC GGGCGGCAGC GCCGCGTCGT CTGGCGTCC CGGGGCGAAG GTCGTGGGG TAGTCCAGGG CTTGGGGTTC GTGGGATGAG TCCGTTGGTG CCGACACCGG GGTTCGGCGA GTGGGGACCG GTGGTGGACG GGACCAGCTG CTAGGGCGAC AACCGCCCGT GCAATCAGGT GAGCTCCCTA GGCAGGCTAG CGCAACAGCT CGGGGCGGGC CGCGGCGCC ATAATGTTGA AAGACTAGGC	CTTGGCCAAGGCCTGATCGGAGCGCTTGTCGCGCACGCCGTCGTGGATACATTGCGAACGATGGTGTCCACATCGCGGTTCTCCAGCGCGTTGAGGTATCGGTTTTGGCCGGTCCCTCCGAGAATGTGCCTGCCGTGTTGGCTCCGTTGGGTATATGATCGCCGCCGTCATAGCCGACACCAGCGCGAGGGCTACCACAACAGCCGCTTGTCGGGTAGGACACCTGCGGCGGCACGCCGGGGGCGGCAGCGCCGCTCGTCTGCCGGTCCGGGGCGAAGGCCGGTTCGGGTCGTGGGGTAGTCCAGGGCTTGGGGTTGTGGGATGAGGGCATTGTGGGTCGTTGGTGCCGACACCGGGTTCGGCGAAACCGCCCGTCGCGTCAGCCGCAATCAGGTGACCACCTAGCCAGCTAGCGCAACAGCTGCCGTCAGCTCGGGGCGGGCCGCGGCCCGATAATGTTGAAAGACTAGGCAACCTTAGGAGAGATTTTGTGACGATCATAATGTTGAAAGACTAGGCAACCTTAGGA

(2) INFORMATION FOR SEQ ID NO:36:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 181 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:36:

GCGGTGTCGG	CGGATCCGGC	GGGTGGTTGA	ACGGCAACGG	CGGGGCCGGC	GGGCCGGCG	60
GGACCGGCGC	TAACGGTGGT	GCCGGCGGCA	ACGCCTGGTT	GTTCGGGGCC	GGCGGGTCCG	120
GCGGNGCCGG	CACCAATGGT	GGNGTCGGCG	GGTCCGGCGG	ATTTGTCTAC	GGCAACGGCG	180
G						181

(2) INFORMATION FOR SEQ ID NO:37:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 290 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:37:

(i) SEQUENCE CHARACTERISTICS:

GCGGTGTCGG CGGATCCGGC GGGTGGTTGA ACGGCAACGG CGGTGTCGGC GGCCGGGGCG	60
GCGACGGCGT CTTTGCCGGT GCCGGCGGCC AGGGCGGCCT CGGTGGGCAG GGCGGCAATG	120
GCGGCGGCTC CACCGGCGGC AACGGCGGTC TTGGCGGCGC GGGCGGTGGC GGAGGCAACG	180
CCCCGGACGG CGGCTTCGGT GGCAACGGCG GTAAGGGTGG CCAGGGCGGN ATTGGCGGCG	240
GCACTCAGAG CGCGACCGGC CTCGGNGGTG ACGGCGGTGA CGGCGGTGAC	290
(2) INFORMATION FOR SEQ ID NO:38:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 34 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:38: GATCCAGTGG CATGGNGGGT GTCAGTGGAA GCAT	34
(2) INFORMATION FOR SEQ ID NO:39:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 155 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:39:	
GATCGCTGCT CGTCCCCCC TTGCCGCCGA CGCCACCGGT CCCACCGTTA CCGAACAAGC	60
TGGCGTGGTC GCCAGCACCC CCGGCACCGC CGACGCCGGA GTCGAACAAT GGCACCGTCG	120
	155
TATCCCCACC ATTGCCGCCG GNCCCACCGG CACCG	135
(2) INFORMATION FOR SEQ ID NO:40:	

(A) LENGTH: 53 base pairs

(B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:40:	
ATGGCGTTCA CGGGGCCCCG GGGACCGGGC AGCCCGGNGG GGCCGGGGGG TGG	53
(2) INFORMATION FOR SEQ ID NO:41:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 132 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:41:	
GATCCACCGC GGGTGCAGAC GGTGCCCGCG GCGCCACCCC GACCAGCGGC GGCAACGGCG	60
GCACCGGCGG CAACGCCGCG AACGCCACCG TCGTCGGNGG GGCCGGCGGG GCCGGCGGCA	120
AGGGCGGCAA CG	132
(2) INFORMATION FOR SEQ ID NO:42:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 132 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:42:	
GATCGGCGGC CGGNACGGNC GGGGACGGCG GCAAGGGCGG NAACGGGGGC GCCGNAGCCA	60
CCNGCCAAGA ATCCTCCGNG TCCNCCAATG GCGCGAATGG CGGACAGGGC GGCAACGGCG	120
GCANCGGCGG CA	132
(2) INFORMATION FOR SEQ ID NO:43:	

(i)	SEQUENCE CHARACTERISTICS:						
	(A)	LENGTH:	702	base	pairs		
			1	:	د:		

(B) TYPE: nucleic acid(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:43:

CGGCACGAGG ATCGGTACCC CGCGGCATCG GCAGCTGCCG ATTCGCCGGG TTTCCCCACC 60 CGAGGAAAGC CGCTACCAGA TGGCGCTGCC GAAGTAGGGC GATCCGTTCG CGATGCCGGC 120 ATGAACGGC GGCATCAAAT TAGTGCAGGA ACCTTTCAGT TTAGCGACGA TAATGGCTAT 180 AGCACTAAGG AGGATGATCC GATATGACGC AGTCGCAGAC CGTGACGGTG GATCAGCAAG 240 AGATTTTGAA CAGGGCCAAC GAGGTGGAGG CCCCGATGGC GGACCCACCG ACTGATGTCC 300 CCATCACACC GTGCGAACTC ACGGNGGNTA AAAACGCCGC CCAACAGNTG GTNTTGTCCG 360 CCGACAACAT GCGGGAATAC CTGGCGGCCG GTGCCAAAGA GCGGCAGCGT CTGGCGACCT 420 CGCTGCGCAA CGCGGCCAAG GNGTATGGCG AGGTTGATGA GGAGGCTGCG ACCGCGCTGG 480 ACAACGACGG CGAAGGAACT GTGCAGGCAG AATCGGCCGG GGCCGTCGGA GGGGACAGTT 540 CGGCCGAACT AACCGATACG CCGAGGGTGG CCACGGCCGG TGAACCCAAC TTCATGGATC 600 TCAAAGAAGC GGCAAGGAAG CTCGAAACGG GCGACCAAGG CGCATCGCTC GCGCACTGNG 660 702 GGGATGGGTG GAACACTINC ACCCTGACGC TGCAAGGCGA CG

(2) INFORMATION FOR SEQ ID NO:44:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 298 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:44:

GAAGCCGCAG CGCTGTCGGG CGACGTGGCG GTCAAAGCGG CATCGCTCGG TGGCGGTGGA 60

GGCGGCGGGG TGCCGTCGGC GCCGTTGGGA TCCGCGATCG GGGCCGCGA ATCGGTGCGG 120

CCCGCTGGCG CTGGTGACAT TGCCGGCTTA GGCCAGGGAA GGGCCGGCGG CGGCGCCGCG 180

CTGGGCGGCG GTGGCATGGG AATGCCGATG GGTGCCGCGC ATCAGGGACA AGGGGGCGCC 240

AAGTCCAAGG GTTCTCAGCA GGAAGACGAG GCGCTCTACA CCGAGGATCC TCGTGCCG 298

(2) INFORMATION FOR SEQ ID NO:45:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1058 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SF	EQUENCE DESC	CRIPTION: SE	EQ ID NO:45:			
CGGCACGAGG	ATCGAATCGC	GTCGCCGGGA	GCACAGCGTC	GCACTGCACC	AGTGGAGGAG	60
CCATGACCTA	CTCGCCGGGT	AACCCCGGAT	ACCCGCAAGC	GCAGCCCGCA	GGCTCCTACG	120
GAGGCGTCAC	ACCCTCGTTC	GCCCACGCCG	ATGAGGGTGC	GAGCAAGCTA	CCGATGTACC	180
TGAACATCGC	GGTGGCAGTG	CTCGGTCTGG	CTGCGTACTT	CGCCAGCTTC	GGCCCAATGT	240
TCACCCTCAG	TACCGAACTC	GGGGGGGTG	ATGGCGCAGT	GTCCGGTGAC	ACTGGGCTGC	300
CGGTCGGGGT	GGCTCTGCTG	GCTGCGCTGC	TTGCCGGGGT	GGTTCTGGTG	CCTAAGGCCA	360
AGAGCCATGT	GACGGTAGTT	GCGGTGCTCG	GGGTACTCGG	CGTATTTCTG	ATGGTCTCGG	420
CGACGTTTAA	CAAGCCCAGC	GCCTATTCGA	CCGGTTGGGC	ATTGTGGGTT	GTGTTGGCTT	480
TCATCGTGTT	CCAGGCGGTT	GCGGCAGTCC	TGGCGCTCTT	GGTGGAGACC	GGCGCTATCA	540
CCGCGCCGGC	GCCGCGGCCC	AAGTTCGACC	CGTATGGACA	GTACGGGCGG	TACGGGCAGT	600
ACGGGCAGTA	CGGGGTGCAG	CCGGGTGGGT	ACTACGGTCA	GCAGGGTGCT	CAGCAGGCCG	660
CGGGACTGCA	GTCGCCCGGC	CCGCAGCAGT	CTCCGCAGCC	TCCCGGATAT	GGGTCGCAGT	720
ACGGCGGCTA	TTCGTCCAGT	CCGAGCCAAT	CGGGCAGTGG	ATACACTGCT	CAGCCCCCGG	780
CCCAGCCGCC	GGCGCAGTCC	GGGTCGCAAC	AATCGCACCA	GGGCCCATCC	ACGCCACCTA	840
CCGGCTTTCC	GAGCTTCAGC	CCACCACCAC	CGGTCAGTGC	CGGGACGGGG	TCGCAGGCTG	900
GTTCGGCTCC	AGTCAACTAT	TCAAACCCCA	GCGGGGGCGA	GCAGTCGTCG	TCCCCGGGG	960
GGGCGCCGGT	CTAACCGGGC	GTTCCCGCGT	CCGGTCGCGC	GTGTGCGCGA	AGAGTGAACA	1020
GGGTGTCAGC	AAGCGCGGAC	GATCCTCGTG	CCGAATTC			1058

(2)	INFORMATION	FOR	SEQ	ID	NO:46:
-----	-------------	-----	-----	----	--------

123	CECHENCE	CHARACTERISTICS:
	SECULIANCE	CHARACIDITE TOO.

- (A) LENGTH: 327 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:46:

CGGCACGAGA	GACCGATGCC	GCTACCCTCG	CGCAGGAGGC	AGGTAATTTC	GAGCGGATCT	60
CCGGCGACCT	GAAAACCCAG	ATCGACCAGG	TGGAGTCGAC	GGCAGGTTCG	TTGCAGGGCC	120
AGTGGCGCGG	CGCGGCGGG	ACGGCCGCCC	AGGCCGCGGT	GGTGCGCTTC	CAAGAAGCAG	180
CCAATAAGCA	GAAGCAGGAA	CTCGACGAGA	TCTCGACGAA	TATTCGTCAG	GCCGGCGTCC	240
AATACTCGAG	GGCCGACGAG	GAGCAGCAGC	AGGCGCTGTC	CTCGCAAATG	GGCTTCTGAC	300
CCGCTAATAC	GAAAAGAAAC	GGAGCAA				327

(2) INFORMATION FOR SEQ ID NO:47:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 170 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:47:

CCAACAACGT GATGGCGTCG TCGAACGTGA CCGATTCTGT ACCGCCGTCG TTGAGATCAA 60

CCAACAACGT GTTGGCGTCG GCAAATGTGC CGNACCCGTG GATCTCGGTG ATCTTGTTCT 120

TCTTCATCAG GAAGTGCACA CCGGCCACCC TGCCCTCGGN TACCTTTCGG 170

(2) INFORMATION FOR SEQ ID NO:48:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 127 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:48:	
GATCCGGCGG CACGGGGGGT GCCGGCGGCA GCACCGCTGG CGCTGGCGGC AACGGCGGGG	60
CCGGGGGTGG CGGCGGAACC GGTGGGTTGC TCTTCGGCAA CGGCGGTGCC GGCGGGCACG	120
GGGCCGT	127
(2) INFORMATION FOR SEQ ID NO:49:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 81 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:49:	
CGGCGGCAAG GGCGGCACCG CCGGCAACGG GAGCGGCGCG GCCGGCGGCA ACGGCGGCAA	60
CGGCGGCTCC GGCCTCAACG G	81
(2) INFORMATION FOR SEQ ID NO:50:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 149 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:50:	
GATCAGGGCT GGCCGGCTCC GGCCAGAAGG GCGGTAACGG AGGAGCTGCC GGATTGTTTG	60
GCAACGGCGG GGCCGGNGGT GCCGGCGCGT CCAACCAAGC CGGTAACGGC GGNGCCGGCG	120
GAAACGGTGG TGCCGGTGGG CTGATCTGG	149
(2) INFORMATION FOR SEQ ID NO:51:	
(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 355 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: single	

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:51:

CGGCACGAGA TCACACCTAC CGAGTGATCG AGATCGTCGG GACCTCGCCC GACGGTGTCG

ACGCGGNAAT CCAGGGCGGT CTGGCCCGAG CTGCGCAGAC CATGCGCGCG CTGGACTGGT

TCGAAGTACA GTCAATTCGA GGCCACCTGG TCGACGGAGC GGTCGCGCAC TTCCAGGTGA

CTATGAAAGT CGGCTTCCGC CTGGAGGATT CCTGAACCTT CAAGCGCGGC CGATAACTGA

GGTGCATCAT TAAGCGACTT TTCCAGAACA TCCTGACGCG CTCGAAACGC GGTTCAGCCG

ACGGTGGCTC CGCCGAGGCG CTGCCTCCAA AATCCCTGCG ACAATTCGTC GGCGG

(2) INFORMATION FOR SEQ ID NO:52:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 999 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:52:

ATGCATCACC ATCACCATCA CATGCATCAG GTGGACCCCA ACTTGACACG TCGCAAGGGA 60 CGATTGGCGG CACTGGCTAT CGCGGCGATG GCCAGCGCCA GCCTGGTGAC CGTTGCGGTG 120 CCCGCGACCG CCAACGCCGA TCCGGAGCCA GCGCCCCCGG TACCCACAAC GGCCGCCTCG 180 CCGCCGTCGA CCGCTGCAGC GCCACCGCA CCGGCGACAC CTGTTGCCCC CCCACCACCG 240 GCCGCCGCCA ACACGCCGAA TGCCCAGCCG GGCGATCCCA ACGCAGCACC TCCGCCGGCC 300 GACCCGAACG CACCGCCGC ACCTGTCATT GCCCCAAACG CACCCCAACC TGTCCGGATC 360 GACAACCCGG TTGGAGGATT CAGCTTCGCG CTGCCTGCTG GCTGGGTGGA GTCTGACGCC 420 GCCCACTTCG ACTACGGTTC AGCACTCCTC AGCAAAACCA CCGGGGACCC GCCATTTCCC 480 GGACAGCCGC CGCCGGTGGC CAATGACACC CGTATCGTGC TCGGCCGGCT AGACCAAAAG 540 CTTTACGCCA GCGCCGAAGC CACCGACTCC AAGGCCGCGG CCCGGTTGGG CTCGGACATG 600 GGTGAGTTCT ATATGCCCTA CCCGGGCACC CGGATCAACC AGGAAACCGT CTCGCTCGAC 660

GCCAACGGGG	TGTCTGGAAG	CGCGTCGTAT	TACGAAGTCA	AGTTCAGCGA	TCCGAGTAAG	720
CCGAACGGCC	AGATCTGGAC	GGGCGTAATC	GGCTCGCCCG	CGGCGAACGC	ACCGGACGCC	780
GGGCCCCCTC	AGCGCTGGTT	TGTGGTATGG	CTCGGGACCG	CCAACAACCC	GGTGGACAAG	840
GGCGCGGCCA	AGGCGCTGGC	CGAATCGATC	CGGCCTTTGG	TCGCCCCGCC	GCCGGCGCCG	900
GCACCGGCTC	CTGCAGAGCC	CGCTCCGGCG	CCGGCGCCGG	CĆGGGGAAGT	CGCTCCTACC	960
CCGACGACAC	CGACACCGCA	GCGGACCTTA	CCGGCCTGA			999

(2) INFORMATION FOR SEQ ID NO:53:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 332 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:53:

Met His His His His His Met His Gln Val Asp Pro Asn Leu Thr
1 5 10 15

Arg Arg Lys Gly Arg Leu Ala Ala Leu Ala Ile Ala Ala Met Ala Ser

Ala Ser Leu Val Thr Val Ala Val Pro Ala Thr Ala Asn Ala Asp Pro 35 40 45

Glu Pro Ala Pro Pro Val Pro Thr Thr Ala Ala Ser Pro Pro Ser Thr 50 55 60

Ala Ala Ala Pro Pro Ala Pro Ala Thr Pro Val Ala Pro Pro Pro 65 70 75 80

Ala Ala Asn Thr Pro Asn Ala Gln Pro Gly Asp Pro Asn Ala Ala 85 90 95

Pro Pro Pro Ala Asp Pro Asn Ala Pro Pro Pro Pro Val Ile Ala Pro 100 105 110

Asn Ala Pro Gln Pro Val Arg Ile Asp Asn Pro Val Gly Gly Phe Ser 115 120 125

Phe Ala Leu Pro Ala Gly Trp Val Glu Ser Asp Ala Ala His Phe Asp 130 135 140

Tyr Gly Ser Ala Leu Leu Ser Lys Thr Thr Gly Asp Pro Pro Phe Pro

145	,				15	0				15	5				160
Gly	Glı	n Pr	o Pr	o Pro 16	Vai	l Ala	a Asr	a Asp	Th:	r Arg	g Ile	e Val	Lev	Gly 175	Arg
Leu	Asp	Gl:	n Ly: 180	s Lei	туз	Ala	Ser	Ala 185	Glu	ı Ala	Thr	Asp	Ser 190		Ala
Ala	Ala	195	J Lei	ı Gly	Ser	Asp	Met 200	Gly	Glu	Phe	Tyr	Met 205	Pro	Tyr	Pro
Gly	Thr 210	Arç	, Il∈	a Asn	Gln	Glu 215	Thr	Val	Ser	Leu	Asp 220	Ala	Asn	Gly	Val
Ser 225	Gly	Ser	Ala	Ser	Tyr 230	Tyr	Glu	Val	Lys	Phe 235	Ser	Asp	Pro	Ser	Lys 240
Pro	Asn	Gly	Gln	Ile 245	Trp	Thr	Gly	Val	Ile 250	Gly	Ser	Pro	Ala	Ala 255	Asn
Ala	Pro	Asp	Ala 260	Gly	Pro	Pro	Gln	Arg 265	Trp	Phe	Val	Val	Trp 270	Leu	Gly
Thr	Ala	Asn 275	Asn	Pro	Val	Asp	Lys 280	Gly	Ala	Ala	Lys	Ala 285	Leu	Ala	Glu
Ser	Ile 290	Arg	Pro	Leu	Val	Ala 295	Pro	Pro	Pro	Ala	Pro 300	Ala	Pro	Ala	Pro
Ala (305	Glu	Pro	Ala	Pro	Ala 310	Pro	Ala	Pro	Ala	Gly 315	Glu	Val	Ala		Thr 320
Pro T	Thr '	Thr	Pro	Thr 325	Pro	Gln .	Arg		Leu 330	Pro	Ala				

(2) INFORMATION FOR SEQ ID NO:54:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 20 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:54:

Asp Pro Val Asp Ala Val Ile Asn Thr Thr Xaa Asn Tyr Gly Gln Val

Val Ala Ala Leu 20

- (2) INFORMATION FOR SEQ ID NO:55:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 15 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55:

Ala Val Glu Ser Gly Met Leu Ala Leu Gly Thr Pro Ala Pro Ser

- (2) INFORMATION FOR SEQ ID NO:56:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:56:

Ala Ala Met Lys Pro Arg Thr Gly Asp Gly Pro Leu Glu Ala Ala Lys

Glu Gly Arg

- (2) INFORMATION FOR SEQ ID NO:57:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 15 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:57:

Tyr Tyr Trp Cys Pro Gly Gln Pro Phe Asp Pro Ala Trp Gly Pro 1

(2) INFORMATION FOR SEQ ID NO:58:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 14 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:58:

Asp Ile Gly Ser Glu Ser Thr Glu Asp Gln Gln Xaa Ala Val 1 5 10

- (2) INFORMATION FOR SEQ ID NO:59:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 13 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:59:

Ala Glu Glu Ser Ile Ser Thr Xaa Glu Xaa Ile Val Pro 1 5 10

- (2) INFORMATION FOR SEQ ID NO:60:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 17 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:60:

Asp Pro Glu Pro Ala Pro Pro Val Pro Thr Ala Ala Ala Ala Pro Pro 1 5 10 15

Ala

(2) INFORMATION FOR SEQ ID NO:61:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 15 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:61:

Ala Pro Lys Thr Tyr Xaa Glu Glu Leu Lys Gly Thr Asp Thr Gly
1 5 10 15

- (2) INFORMATION FOR SEQ ID NO:62:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 30 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:62:

Asp Pro Ala Ser Ala Pro Asp Val Pro Thr Ala Ala Gln Gln Thr Ser 1 5 10 15

Leu Leu Asn Asn Leu Ala Asp Pro Asp Val Ser Phe Ala Asp 20 25 30

- (2) INFORMATION FOR SEQ ID NO:63:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 24 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:63:

Gly Cys Gly Asp Arg Ser Gly Gly Asn Leu Asp Gln Ile Arg Leu Arg 1 5 10 15

Arg Asp Arg Ser Gly Gly Asn Leu 20

(2) INFORMATION FOR SEQ ID NO:64:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 187 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:64:

Thr Gly Ser Leu Asn Gln Thr His Asn Arg Arg Ala Asn Glu Arg Lys
1 10 15

Asn Thr Thr Met Lys Met Val Lys Ser Ile Ala Ala Gly Leu Thr Ala 20 25 30

Ala Ala Ala Ile Gly Ala Ala Ala Ala Gly Val Thr Ser Ile Met Ala 35 . 40 45

Gly Gly Pro Val Val Tyr Gln Met Gln Pro Val Val Phe Gly Ala Pro 50 55 60

Leu Pro Leu Asp Pro Ala Ser Ala Pro Asp Val Pro Thr Ala Ala Gln 65 70 75 80

Leu Thr Ser Leu Leu Asn Ser Leu Ala Asp Pro Asn Val Ser Phe Ala 85 90 95

Asn Lys Gly Ser Leu Val Glu Gly Gly Ile Gly Gly Thr Glu Ala Arg 100 105 110

Ile Ala Asp His Lys Leu Lys Lys Ala Ala Glu His Gly Asp Leu Pro 115 120 125

Leu Ser Phe Ser Val Thr Asn Ile Gln Pro Ala Ala Ala Gly Ser Ala 130 135 140

Thr Ala Asp Val Ser Val Ser Gly Pro Lys Leu Ser Ser Pro Val Thr 145 150 155 160

Gln Asn Val Thr Phe Val Asn Gln Gly Gly Trp Met Leu Ser Arg Ala 165 170 175

Ser Ala Met Glu Leu Leu Gln Ala Ala Gly Xaa 180 185

(2) INFORMATION FOR SEQ ID NO:65:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 148 amino acids
 - (B) TYPE: amino acid

- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:65:

Asp Glu Val Thr Val Glu Thr Thr Ser Val Phe Arg Ala Asp Phe Leu 1 5 10 15

Ser Glu Leu Asp Ala Pro Ala Gln Ala Gly Thr Glu Ser Ala Val Ser 20 25 30

Gly Val Glu Gly Leu Pro Pro Gly Ser Ala Leu Leu Val Val Lys Arg 35 40 45

Gly Pro Asn Ala Gly Ser Arg Phe Leu Leu Asp Gln Ala Ile Thr Ser 50 60

Ala Gly Arg His Pro Asp Ser Asp Ile Phe Leu Asp Asp Val Thr Val 65 70 75 80

Ser Arg Arg His Ala Glu Phe Arg Leu Glu Asn Asn Glu Phe Asn Val 85 90 95

Val Asp Val Gly Ser Leu Asn Gly Thr Tyr Val Asn Arg Glu Pro Val 100 105 110

Asp Ser Ala Val Leu Ala Asn Gly Asp Glu Val Gln Ile Gly Lys Leu 115 120 125

Arg Leu Val Phe Leu Thr Gly Pro Lys Gln Gly Glu Asp Asp Gly Ser 130 135 140

Thr Gly Gly Pro 145

- (2) INFORMATION FOR SEQ ID NO:66:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 230 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:66:

Thr Ser Asn Arg Pro Ala Arg Arg Gly Arg Arg Ala Pro Arg Asp Thr 1 5 10 15

- Gly Pro Asp Arg Ser Ala Ser Leu Ser Leu Val Arg His Arg Arg Gln
 20 25 30
- Gln Arg Asp Ala Leu Cys Leu Ser Ser Thr Gln Ile Ser Arg Gln Ser 35 40 45
- Asn Leu Pro Pro Ala Ala Gly Gly Ala Ala Asn Tyr Ser Arg Asn 50 55 60
- Phe Asp Val Arg Ile Lys Ile Phe Met Leu Val Thr Ala Val Val Leu 65 70 75 80
- Leu Cys Cys Ser Gly Val Ala Thr Ala Ala Pro Lys Thr Tyr Cys Glu 85 90
- Glu Leu Lys Gly Thr Asp Thr Gly Gln Ala Cys Gln Ile Gln Met Ser
- Asp Pro Ala Tyr Asn Ile Asn Ile Ser Leu Pro Ser Tyr Tyr Pro Asp 115 120 125
- Gln Lys Ser Leu Glu Asn Tyr Ile Ala Gln Thr Arg Asp Lys Phe Leu 130 135 140
- Ser Ala Ala Thr Ser Ser Thr Pro Arg Glu Ala Pro Tyr Glu Leu Asn 145 150 155 160
- Ile Thr Ser Ala Thr Tyr Gln Ser Ala Ile Pro Pro Arg Gly Thr Gln
 165 170
- Ala Val Val Leu Xaa Val Tyr His Asn Ala Gly Gly Thr His Pro Thr 180 185 190
- Thr Thr Tyr Lys Ala Phe Asp Trp Asp Gln Ala Tyr Arg Lys Pro Ile
 195 200 205
- Thr Tyr Asp Thr Leu Trp Gln Ala Asp Thr Asp Pro Leu Pro Val Val 210 215 220

Phe Pro Ile Val Ala Arg 225 230

(2) INFORMATION FOR SEQ ID NO:67:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 132 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:67:

Thr Ala Ala Ser Asp Asn Phe Gln Leu Ser Gln Gly Gln Gly Phe
1 5 10 15

Ala Ile Pro Ile Gly Gln Ala Met Ala Ile Ala Gly Gln Ile Arg Ser 20 25 30

Gly Gly Gly Ser Pro Thr Val His Ile Gly Pro Thr Ala Phe Leu Gly 35 40 45

Leu Gly Val Val Asp Asn Asn Gly Asn Gly Ala Arg Val Gln Arg Val 50 55 60

Val Gly Ser Ala Pro Ala Ala Ser Leu Gly Ile Ser Thr Gly Asp Val 65 70 75 80

Ile Thr Ala Val Asp Gly Ala Pro Ile Asn Ser Ala Thr Ala Met Ala 85 90 95

Asp Ala Leu Asn Gly His His Pro Gly Asp Val Ile Ser Val Asn Trp 100 105 110

Gln Thr Lys Ser Gly Gly Thr Arg Thr Gly Asn Val Thr Leu Ala Glu 115 120 125

Gly Pro Pro Ala 130

- (2) INFORMATION FOR SEQ ID NO:68:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 100 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:68:

Val Pro Leu Arg Ser Pro Ser Met Ser Pro Ser Lys Cys Leu Ala Ala 1 5 10 15

Ala Gln Arg Asn Pro Val Ile Arg Arg Arg Arg Leu Ser Asn Pro Pro 20 25 30

Pro Arg Lys Tyr Arg Ser Met Pro Ser Pro Ala Thr Ala Ser Ala Gly 35 40 45

Met Ala Arg Val Arg Arg Ala Ile Trp Arg Gly Pro Ala Thr Xaa 50 55 60

Ser Ala Gly Met Ala Arg Val Arg Arg Trp Xaa Val Met Pro Xaa Val 65 70 75 80

Ile Gln Ser Thr Xaa Ile Arg Xaa Xaa Gly Pro Phe Asp Asn Arg Gly 85 90 95

Ser Glu Arg Lys 100

(2) INFORMATION FOR SEQ ID NO:69:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 163 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:69:

Met Thr Asp Asp Ile Leu Leu Ile Asp Thr Asp Glu Arg Val Arg Thr 1 5 10 15

Leu Thr Leu Asn Arg Pro Gln Ser Arg Asn Ala Leu Ser Ala Ala Leu 20 25 30

Arg Asp Arg Phe Phe Ala Xaa Leu Xaa Asp Ala Glu Xaa Asp Asp Asp Asp 35

Ile Asp Val Val Ile Leu Thr Gly Ala Asp Pro Val Phe Cys Ala Gly 50 55 60

Leu Asp Leu Lys Val Ala Gly Arg Ala Asp Arg Ala Ala Gly His Leu 65 70 75 80

Thr Ala Val Gly Gly His Asp Gln Ala Gly Asp Arg Arg Asp Gln Arg 85 90 95

Arg Arg Gly His Arg Arg Ala Arg Thr Gly Ala Val Leu Arg His Pro
100 105 110

Asp Arg Leu Arg Ala Arg Pro Leu Arg Arg His Pro Arg Pro Gly Gly 115 120 125

Ala Ala His Leu Gly Thr Gln Cys Val Leu Ala Ala Lys Gly Arg 130 135 140

His Arg Xaa Gly Pro Val Asp Glu Pro Asp Arg Arg Leu Pro Val Arg 145 150 155 160

Asp Arg Arg

(2) INFORMATION FOR SEQ ID NO:70:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 344 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:70: Met Lys Phe Val Asn His Ile Glu Pro Val Ala Pro Arg Arg Ala Gly 10 Gly Ala Val Ala Glu Val Tyr Ala Glu Ala Arg Arg Glu Phe Gly Arg Leu Pro Glu Pro Leu Ala Met Leu Ser Pro Asp Glu Gly Leu Leu Thr 40 Ala Gly Trp Ala Thr Leu Arg Glu Thr Leu Leu Val Gly Gln Val Pro Arg Gly Arg Lys Glu Ala Val Ala Ala Val Ala Ala Ser Leu Arg 70 Cys Pro Trp Cys Val Asp Ala His Thr Thr Met Leu Tyr Ala Ala Gly 90 Gln Thr Asp Thr Ala Ala Ala Ile Leu Ala Gly Thr Ala Pro Ala Ala 105 Gly Asp Pro Asn Ala Pro Tyr Val Ala Trp Ala Ala Gly Thr Gly Thr 120 Pro Ala Gly Pro Pro Ala Pro Phe Gly Pro Asp Val Ala Ala Glu Tyr 135 Leu Gly Thr Ala Val Gln Phe His Phe Ile Ala Arg Leu Val Leu Val 150 145 Leu Leu Asp Glu Thr Phe Leu Pro Gly Gly Pro Arg Ala Gln Gln Leu 170

Met Arg Arg Ala Gly Gly Leu Val Phe Ala Arg Lys Val Arg Ala Glu 185

His Arg Pro Gly Arg Ser Thr Arg Arg Leu Glu Pro Arg Thr Leu Pro

Asp Asp Leu Ala Trp Ala Thr Pro Ser Glu Pro Ile Ala Thr Ala Phe

200

195

104

210 215 220

Ala Ala Leu Ser His His Leu Asp Thr Ala Pro His Leu Pro Pro Pro 225 230 235 240

Thr Arg Gln Val Val Arg Arg Val Val Gly Ser Trp His Gly Glu Pro
245 250 255

Met Pro Met Ser Ser Arg Trp Thr Asn Glu His Thr Ala Glu Leu Pro 260 265 270

Ala Asp Leu His Ala Pro Thr Arg Leu Ala Leu Leu Thr Gly Leu Ala 275 280 285

Pro His Gln Val Thr Asp Asp Val Ala Ala Ala Arg Ser Leu Leu 290 295 300

Asp Thr Asp Ala Ala Leu Val Gly Ala Leu Ala Trp Ala Ala Phe Thr 305 310 315 / 320

Ala Ala Arg Arg Ile Gly Thr Trp Ile Gly Ala Ala Ala Glu Gly Gln
325 330 335

Val Ser Arg Gln Asn Pro Thr Gly 340

(2) INFORMATION FOR SEQ ID NO:71:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 485 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:71:

Asp Asp Pro Asp Met Pro Gly Thr Val Ala Lys Ala Val Ala Asp Ala 1 5 10 15

Leu Gly Arg Gly Ile Ala Pro Val Glu Asp Ile Gln Asp Cys Val Glu 20 25 30

Ala Arg Leu Gly Glu Ala Gly Leu Asp Asp Val Ala Arg Val Tyr Ile 35 40 45

Ile Tyr Arg Gln Arg Arg Ala Glu Leu Arg Thr Ala Lys Ala Leu Leu 50 55 60

Gly Val Arg Asp Glu Leu Lys Leu Ser Leu Ala Ala Val Thr Val Leu 65 70 75 80

Arg	Glu	Arg	Tyr	Leu 85	Leu	His	Asp	Glu	Gln 90	Gly	Arg	Pro	Ala	Glu 95	Ser
Thr	Gly	Glu	Leu 100	Met	Asp	Arg	Ser	Ala 105	Arg	Cys	Val	Ala	Ala 110	Ala	Glu
Asp	Gln	Tyr 115	Glu	Pro	Gly	Ser	Ser 120	Arg	Arg	Trp	Ala	Glu 125	Arg	Phe	Ala
Thr	Leu 130	Leu	Arg	Asn	Leu	Glu 135	Phe	Leu	Pro	Asn	Ser 140	Pro	Thr	Leu	Met
Asn 145	Ser	Gly	Thr	Asp	Leu 150	Gly	Leu	Leu	Ala	Gly 155	Суѕ	Phe	Val	Leu	Pro 160
Ile	Glu	Asp	Ser	Leu 165	Gln	Ser	Ile	Phe	Ala 170	Thr	Leu	Gly	Gln	Ala 175 /	Ala
Glu	Leu	Gln	Arg 180	Ala	Gly	Gly	Gly	Thr 185	Gly	Tyr	Ala	Phe	Ser 190	His	Leu
Arg	Pro	Ala 195	Gly	Asp	Arg	Val	Ala 200	Ser	Thr	Gly	Gly	Thr 205	Ala	Ser	Gly
Pro	Val 210	Ser	Phe	Leu	Arg	Leu 215	Tyr	Asp	Ser	Ala	Ala 220	Gly	Val	Val	Ser
Met 225	Gly	Gly	Arg	Arg	Arg 230	Gly	Ala	Cys	Met	Ala 235	Val	Leu	Asp	Val	Ser 240
His	Pro	Asp	Ile	Cys 245	Asp	Phe	Val	Thr	Ala 250	Lys	Ala	Glu	Ser	Pro 255	Ser
Glu	Leu	Pro	His 260	Phe	Asn	Leu	Ser	Val 265	Gly	Val	Thr	Asp	Ala 270	Phe	Leu
Arg	Ala	Val 275	Glu	Arg	Asn	Gly	Leu 280	His	Arg	Leu	Val	Asn 285	Pro	Arg	Thr
Gly	Lys 290	Ile	Val	Ala	Arg	Met 295	Pro	Ala	Ala	Glu	Leu 300	Phe	Asp	Ala	Ile
Cys 305	Lys	Ala	Ala	His	Ala 310	Gly	Gly	Asp	Pro	Gly 315	Leu	Val	Phe	Leu	Asp 320
Thr	Ile	Asn	Arg	Ala 325	Asn	Pro	Val	Pro	Gly 330	Arg	Gly	Arg	Ile	Glu 335	Ala
Thr	Asn	Pro	Cys 340	Gly	Glu	Val	Pro	Leu 345	Leu	Pro	Tyr	Glu	Ser 350	Cys	Asn
Leu	Gly	Ser 355	Ile	Asn	Leu	Ala	Arg 360	Met	Leu	Ala	Asp	Gly 365		Val	Asp
Trp	Asp	Arg	Leu	Glu	Glu	Val	Ala	Gly	Val	Ala	Val	Arg	Phe	Leu	Asp

106

370 375 380

Asp Val Ile Asp Val Ser Arg Tyr Pro Phe Pro Glu Leu Gly Glu Ala 385 390 395 400

Ala Arg Ala Thr Arg Lys Ile Gly Leu Gly Val Met Gly Leu Ala Glu 405 410 415

Leu Leu Ala Ala Leu Gly Ile Pro Tyr Asp Ser Glu Glu Ala Val Arg 420 425 . 430

Leu Ala Thr Arg Leu Met Arg Arg Ile Gln Gln Ala Ala His Thr Ala 435 440 445

Ser Arg Arg Leu Ala Glu Glu Arg Gly Ala Phe Pro Ala Phe Thr Asp 450 455 460

Ser Arg Phe Ala Arg Ser Gly Pro Arg Arg Asn Ala Gln Val Thr Ser 465 470 475 480

Val Ala Pro Thr Gly 485

(2) INFORMATION FOR SEQ ID NO:72:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 267 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:72:

Gly Val Ile Val Leu Asp Leu Glu Pro Arg Gly Pro Leu Pro Thr Glu
1 10 15

Ile Tyr Trp Arg Arg Gly Leu Ala Leu Gly Ile Ala Val Val 20 25 30

Val Gly Ile Ala Val Ala Ile Val Ile Ala Phe Val Asp Ser Ser Ala 35 40 45

Gly Ala Lys Pro Val Ser Ala Asp Lys Pro Ala Ser Ala Gln Ser His 50 55 60

Pro Gly Ser Pro Ala Pro Gln Ala Pro Gln Pro Ala Gly Gln Thr Glu 65 70 75 80

Gly Asn Ala Ala Ala Pro Pro Gln Gly Gln Asn Pro Glu Thr Pro 85 90 95

- Thr Pro Thr Ala Ala Val Gln Pro Pro Pro Val Leu Lys Glu Gly Asp Asp Cys Pro Asp Ser Thr Leu Ala Val Lys Gly Leu Thr Asn Ala Pro 115 Gln Tyr Tyr Val Gly Asp Gln Pro Lys Phe Thr Met Val Val Thr Asn Ile Gly Leu Val Ser Cys Lys Arg Asp Val Gly Ala Ala Val Leu Ala 150 155 Ala Tyr Val Tyr Ser Leu Asp Asn Lys Arg Leu Trp Ser Asn Leu Asp 170 Cys Ala Pro Ser Asn Glu Thr Leu Val Lys Thr Phe Ser Pro Gly Glu 185 Gln Val Thr Thr Ala Val Thr Trp Thr Gly Met Gly Ser Ala Pro Arg Cys Pro Leu Pro Arg Pro Ala Ile Gly Pro Gly Thr Tyr Asn Leu Val Val Gln Leu Gly Asn Leu Arg Ser Leu Pro Val Pro Phe Ile Leu Asn 235 230 Gln Pro Pro Pro Pro Gly Pro Val Pro Ala Pro Gly Pro Ala Gln 250 245 Ala Pro Pro Pro Glu Ser Pro Ala Gln Gly Gly
- (2) INFORMATION FOR SEQ ID NO:73:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 97 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:73:
 - Leu Ile Ser Thr Gly Lys Ala Ser His Ala Ser Leu Gly Val Gln Val
 - Thr Asn Asp Lys Asp Thr Pro Gly Ala Lys Ile Val Glu Val Val Ala
 20 25 30
 - Gly Gly Ala Ala Asn Ala Gly Val Pro Lys Gly Val Val Val Thr 35 40 45

Lys Val Asp Asp Arg Pro Ile Asn Ser Ala Asp Ala Leu Val Ala Ala 50 55 60

Val Arg Ser Lys Ala Pro Gly Ala Thr Val Ala Leu Thr Phe Gln Asp
65 70 75 80

Pro Ser Gly Gly Ser Arg Thr Val Gln Val Thr Leu Gly Lys Ala Glu 85 90 95

Gln

(2) INFORMATION FOR SEQ ID NO:74:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 364 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:74:
- Gly Ala Ala Val Ser Leu Leu Ala Ala Gly Thr Leu Val Leu Thr Ala 1 5 10 15
- Cys Gly Gly Gly Thr Asn Ser Ser Ser Gly Ala Gly Gly Thr Ser 20 25 30
- Gly Ser Val His Cys Gly Gly Lys Lys Glu Leu His Ser Ser Gly Ser 35 40 45
- Thr Ala Gln Glu Asn Ala Met Glu Gln Phe Val Tyr Ala Tyr Val Arg 50 55 60
- Ser Cys Pro Gly Tyr Thr Leu Asp Tyr Asn Ala Asn Gly Ser Gly Ala 65 70 75 80
- Gly Val Thr Gln Phe Leu Asn Asn Glu Thr Asp Phe Ala Gly Ser Asp 85 90 95
- Val Pro Leu Asn Pro Ser Thr Gly Gln Pro Asp Arg Ser Ala Glu Arg
 100 105 110
- Cys Gly Ser Pro Ala Trp Asp Leu Pro Thr Val Phe Gly Pro Ile Ala 115 120 125
- Ile Thr Tyr Asn Ile Lys Gly Val Ser Thr Leu Asn Leu Asp Gly Pro 130 135 140
- Thr Thr Ala Lys Ile Phe Asn Gly Thr Ile Thr Val Trp Asn Asp Pro

145					150					155					160
Gln	Ile	Gln	Ala	Leu 165	Asn	Ser	Gly	Thr	Asp 170	Leu	Pro	Pro	Thr	Pro 175	Ile
Ser	Val	Ile	Phe 180	Arg	Ser	Asp	Lys	Ser 185	Gly	Thr	Ser	Asp	Asn 190	Phe	Gln
Lys	Tyr	Leu 195	Asp	Gly	Val	Ser	Asn 200	Gly	Ala	Trp	Gly	Lys 205	Gly	Ala	Ser
Glu	Thr 210	Phe	Ser	Gly	Gly	Val 215	Gly	Val	Gly	Ala	Ser 220	Gly	Asn	Asn	Gly
Thr 225	Ser	Ala	Leu	Leu	Gln 230	Thr	Thr	Asp	Gly	Ser 235	Ile	Thr	Tyr	Asn	Glu 240
Trp	Ser	Phe	Ala	Val 245	Gly	Lys	Gln	Leu	Asn 250	Met	Ala	Gln	Ile	Ile 255	Thr
Ser	Ala	Gly	Pro 260	Asp	Pro	Val	Ala	Ile 265	Thr	Thr	Glu	Ser	Val 270	Gly	Lys
Thr	Ile	Ala 275	Gly	Ala	Lys	Ile	Met 280	Gly	Gln	Gly	Asn	Asp 285	Leu	Val	Leu
Asp	Thr 290	Ser	Ser	Phe	Tyr	Arg 295	Pro	Thr	Gln	Pro	Gly 300	Ser	Tyr	Pro	Ile
Val 305	Leu	Ala	Thr	Tyr	Glu 310	Ile	Val	Cys	Ser	Lys 315	Tyr	Pro	Asp	Ala	Thr 320
Thr	Gly	Thr	Ala	Val 325	Arg	Ala	Phe	Met	Gln 330		Ala	Ile	Gly	Pro 335	Gly
Gln	Glu	Gly	Leu 340	Asp	Gln	Tyr	Gly	Ser 345	Ile	Pro	Leu	Pro	350	Ser	Phe
Gln	Ala	Lys 355	Leu	Ala	Ala	Ala	Val 360	Asn	Ala	Ile	Ser	•			

(2) INFORMATION FOR SEQ ID NO:75:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 309 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:75:

Gln 1	Ala	Ala	Ala	Gly 5	Arg	Ala	Val	Arg	Arg 10	Thr	Gly	His	Ala	Glu 15	Asp
Gln	Thr	His	Gln 20	Asp	Arg	Leu	His	His 25	Gly	Cys	Arg	Arg	Ala 30	Ala	Val
Val	Val	Arg 35	Gln	Asp	Arg	Ala	Ser 40	Val	Ser	Ala	Thr	Ser 45	Ala	Arg	Pro
Pro	Arg 50	Arg	His	Pro	Ala	Gln 55	Gly	His	Arg	Arg	Arg 60	Val	Ala	Pro	Ser
Gly 65	Gly	Arg	Arg	Arg	Pro 70	His	Pro	His	His	Val 75	Gln	Pro	Asp	Asp	Arg 80
Arg	Asp	Arg	Pro	Ala 85	Leu	Leu	Asp	Arg	Thr 90	Gln	Pro	Ala	Glu	His 95 /	Pro
Asp	Pro	His	Arg 100	Arg	Gly	Pro	Ala	Asp 105	Pro	Gly	Arg	Val	Arg 110	G1/y	Arg
Gly	Arg	Leu 115	Arg	Arg	Val	Asp	Asp 120	Gly	Arg	Leu	Gln	Pro 125	Asp	Arg	Asp
	130					135					140		Arġ		
145					150					155			Gly		160
-				165					170				Gly	175	
-			180					185					Pro 190		
		195					200					205	Gly		
	210					215					220		Arg		
225	_				230					235			Pro		240
				245					250				Gly	255	
			260			ż		265					Gly 270		
		275					280					285			
707 -	C1	1721	Δla	Hie	בומ	Δla	Δla	Glv	Pro	Ara	Ara	Ala	Ala	Val	Arc

290 295 300

Asn Arg Pro Arg Arg 305

(2) INFORMATION FOR SEQ ID NO:76:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 580 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi)	SEQ	JENC	E DE	SCRI	PTIO	N: SI	EQ II	ONO:	76:						
Ser 1	Ala	Val	Trp	Cys 5	Leu	Asn	Gly	Phe	Thr 10	Gly	Arg	His	Arg	His 15	Gly
Arg	Cys	Arg	Val 20	Arg	Ala	Ser	Gly	Trp 25	Arg	Ser	Ser	Asn	Arg 30	Trp	Cys
Ser	Thr	Thr 35	Ala	Asp	Суѕ	Cys	Ala 40	Ser	Lys	Thr	Pro	Thr 45	Gln	Ala	Ala
Ser	Pro 50	Leu	Glu	Arg	Arg	Phe 55	Thr	Cys	Cys	Ser	Pro 60	Ala	Val	Gly	Cys
Arg 65	Phe	Arg	Ser	Phe	Pro 70	Val	Arg	Arg	Leu	Ala 75	Leu	Gly	Ala	Arg	Thr 80
Ser	Arg	Thr	Leu	Gly 85	Val	Arg	Arg	Thr	Leu 90	Ser	Gln	Trp	Asn	Leu 95	Ser
Pro	Arg	Ala	Gln 100	Pro	Ser	Cys	Ala	Val 105	Thr	Val	Glu	Ser	His 110	Thr	His
Ala	Ser	Pro 115	Arg	Met	Ala	Lys	Leu 120	Ala	Arg	Val	Val	Gly 125	Leu	Val	Gln
Glu	Glu 130	Gln	Pro	Ser	Asp	Met 135	Thr	Asn	His	Pro	Arg 140	Tyr	Ser	Pro	Pro
Pro 145	Gln	Gln	Pro	Gly	Thr 150	Pro	Gly	Tyr	Ala	Gln 155	Gly	Gln	Gln	Gln	Thr 160
Tyr	Ser	Gln	Gln	Phe 165	Asp	Trp	Arg	Tyr	Pro 170	Pro	Ser	Pro	Pro	Pro 175	Gln
Pro	Thr	Gln	Туг 180	Arg	Gln	Pro	Tyr	Glu 185	Ala	Leu	Gly	Gly	Thr 190	Arg	Pro

Gly	Leu	Ile 195	Pro	Gly	Val	Ile	Pro 200	Thr	Met	Thr	Pro	Pro 205	Pro	Gly	Met
Val	Arg 210	Gln	Arg	Pro	Arg	Ala 215	Gly	Met	Leu	Ala	Ile 220	Gly	Ala	Val	Thr
Ile 225	Ala	Val	Val	Ser	Ala 230	Gly	Ile	Gly	Gly	Ala 235	Ala	Ala	Ser	Leu	Val 240
Gly	Phe	Asn	Arg	Ala 245	Pro	Ala	Gly	Pro	Ser 250	Gly	Gly	Pro	Val	Ala 255	Ala
Ser	Ala	Ala	Pro 260	Ser	Ile	Pro	Ala	Ala 265	Asn	Met	Pro	Pro	Gly 270	Ser	Val
Glu	Gln	Val 275	Ala	Ala	Lys	Val	Val 280	Pro	Ser	Val	Val	Met 285	Leu	Glu	Thr
Asp	Leu 290	Gly	Arg	Gln	Ser	Glu 295	Glu	Gly	Ser	Gly	11e 300	Ile	Leu	Ser	Ala
Glu 305	Gly	Leu	Ile	Leu	Thr 310	Asn	Asn	His	Val	11e 315	Ala	Ala	Ala	Ala	Lys 320
Pro	Pro	Leu	Gly	Ser 325	Pro	Pro	Pro	Lys	Thr 330	Thr	Val	Thr	Phe	Ser 335	Asp
Gly	Arg	Thr	Ala 340	Pro	Phe	Thr	Val	Val 345	Gly	Ala	Asp	Pro	Thr 350	Ser	Asp
Ile	Ala	Val 355	Val	Arg	Val	Gln	Gly 360	Val	Ser	Gly	Leu	Thr 365	Pro	Ile	Ser
Leu	Gly 370	Ser	Ser	Ser	Asp	Leu 375	Arg	Val	Gly	Gln	Pro 380	Val	Leu	Ala	Ile
385		•			390					395					Ser 400
Ala	Leu	Asn	Arg	Pro 405	Val	Ser	Thr	Thr	Gly 410		Ala	Gly	Asn	Gln 415	Asn
Thr	Val	Leu	Asp 420	Ala	Ile	Gln	Thr	Asp 425		Ala	Ile	Asn	Pro 430		Asn
Ser	Gly	Gly 435	Ala	Leu	Val	Asn	Met 440		Ala	Gln	Leu	Val 445	Gly	Val	Asn
Ser	Ala 450	Ile	Ala	Thr	Leu	Gly 455		Asp	Ser	Ala	Asp 460		Gln	Ser	Gly
Ser 465		Gly	Leu	Gly	Phe 470	Ala	Ile	Pro	Val	Asp 475		Ala	Lys	Arg	Ile 480
Ala	Asp	Glu	Leu	Ile	Ser	Thr	Gly	Lys	Ala	Ser	His	Ala	Ser	Leu	Gly

485 490 495

Val Gln Val Thr Asn Asp Lys Asp Thr Pro Gly Ala Lys Ile Val Glu 500 505 510

Val Val Ala Gly Gly Ala Ala Ala Asn Ala Gly Val Pro Lys Gly Val 515 520 525

Val Val Thr Lys Val Asp Asp Arg Pro Ile Asn Ser Ala Asp Ala Leu 530 535 540

Val Ala Ala Val Arg Ser Lys Ala Pro Gly Ala Thr Val Ala Leu Thr 545 550 555 560

Phe Gln Asp Pro Ser Gly Gly Ser Arg Thr Val Gln Val Thr Leu Gly 565 570 575

Lys Ala Glu Gln 580

(2) INFORMATION FOR SEQ ID NO:77:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 233 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:77:

Met Asn Asp Gly Lys Arg Ala Val Thr Ser Ala Val Leu Val Val Leu
1 10 15

Gly Ala Cys Leu Ala Leu Trp Leu Ser Gly Cys Ser Ser Pro Lys Pro 20 25 30

Asp Ala Glu Glu Gln Gly Val Pro Val Ser Pro Thr Ala Ser Asp Pro 35 40 45

Ala Leu Leu Ala Glu Ile Arg Gln Ser Leu Asp Ala Thr Lys Gly Leu 50 55 60

Thr Ser Val His Val Ala Val Arg Thr Thr Gly Lys Val Asp Ser Leu 65 70 75 80

Leu Gly Ile Thr Ser Ala Asp Val Asp Val Arg Ala Asn Pro Leu Ala 85 90 95

Ala Lys Gly Val Cys Thr Tyr Asn Asp Glu Gln Gly Val Pro Phe Arg 100 105 110

Val Gln Gly Asp Asn Ile Ser Val Lys Leu Phe Asp Asp Trp Ser Asn 115 120 125

Leu Gly Ser Ile Ser Glu Leu Ser Thr Ser Arg Val Leu Asp Pro Ala 130 135 140

Ala Gly Val Thr Gln Leu Leu Ser Gly Val Thr Asn Leu Gln Ala Gln 145 150 155 . 160

Gly Thr Glu Val Ile Asp Gly Ile Ser Thr Thr Lys Ile Thr Gly Thr 165 170 175

Ile Pro Ala Ser Ser Val Lys Met Leu Asp Pro Gly Ala Lys Ser Ala 180 185 190

Arg Pro Ala Thr Val Trp Ile Ala Gln Asp Gly Ser His His Leu Val 195 200 205

Arg Ala Ser Ile Asp Leu Gly Ser Gly Ser Ile Gln Leu Thr Gln Ser 210 215 220

Lys Trp Asn Glu Pro Val Asn Val Asp 225 230

(2) INFORMATION FOR SEQ ID NO:78:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 66 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:78:

Val Ile Asp Ile Ile Gly Thr Ser Pro Thr Ser Trp Glu Gln Ala Ala 1 5 10 15

Ala Glu Ala Val Gln Arg Ala Arg Asp Ser Val Asp Asp Ile Arg Val 20 25 30

Ala Arg Val Ile Glu Gln Asp Met Ala Val Asp Ser Ala Gly Lys Ile 35 40 45

Thr Tyr Arg Ile Lys Leu Glu Val Ser Phe Lys Met Arg Pro Ala Gln 50 55 60

Pro Arg 65

(2) INFORMATION FOR SEQ ID NO:79:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 69 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:79:

Val Pro Pro Ala Pro Pro Leu Pro Pro Leu Pro Pro Ser Pro Ile Ser 1 5 10 15

Cys Ala Ser Pro Pro Ser Pro Pro Leu Pro Pro Ala Pro Pro Val Ala 20 25 30

Pro Gly Pro Pro Met Pro Pro Leu Asp Pro Trp Pro Pro Ala Pro Pro 35 40 45

Leu Pro Tyr Ser Thr Pro Pro Gly Ala Pro Leu Pro Pro Ser Pro Pro 50 55 60

Ser Pro Pro Leu Pro 65

- (2) INFORMATION FOR SEQ ID NO:80:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 355 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:80:

Met Ser Asn Ser Arg Arg Arg Ser Leu Arg Trp Ser Trp Leu Leu Ser 1 5 10 15

Val Leu Ala Ala Val Gly Leu Gly Leu Ala Thr Ala Pro Ala Gln Ala 20 25 30

Ala Pro Pro Ala Leu Ser Gln Asp Arg Phe Ala Asp Phe Pro Ala Leu 35 40 45

Pro Leu Asp Pro Ser Ala Met Val Ala Gln Val Ala Pro Gln Val Val 50 55 60

Asn Ile Asn Thr Lys Leu Gly Tyr Asn Asn Ala Val Gly Ala Gly Thr 65 70 75 80

Gly	Ile	Val	Ile	Asp 85	Pro	Asn	Gly	Val	Val 90	Leu	Thr	Asn	Asn	His 95	Val
Ile	Ala	Gly	Ala 100	Thr	Asp	Ile	Asn	Ala 105	Phe	Ser	Val	Gly	Ser 110	Gly	Gln
Thr	Tyr	Gly 115	Val	Asp	Val	Val	Gly 120	Tyr	Asp	Arg	Thr	Gln 125	Asp	Val	Ala
Val	Leu 130	Gln	Leu	Arg	Gly	Ala 135	Gly	Gly	Leu	Pro	Ser 140	Ala	Ala	Ile	Gly
Gly 145	Gly	Val	Ala	Val	Gly 150	Glu	Pro	Val	Val	Ala 155	Met	Gly	Asn	Ser	Gly 160
Gly	Gln	Gly	Gly	Thr 165	Pro	Arg	Ala	Val	Pro 170	Gly	Arg	Val	Val	Ala 175	Leu
Gly	Gln	Thr	Val 180	Gln	Ala	Ser	Asp	Ser 185	Leu	Thr	Gly	Ala	Glu 190	Glu	Thr
Leu	Asn	Gly 195	Leu	Ile	Gln	Phe	Asp 200	Ala	Ala	Ile	Gln	Pro 205	Gly	Asp	Ser
Gly	Gly 210	Pro	Val	Val	Asn	Gly 215	Leu	Gly	Gln	Val	Val 220	Gly	Met	Asn	Thr
Ala 225	Ala	Ser	Asp	Asn	Phe 230	Gln	Leu	Ser	Gln	Gly 235	Gly	Gln	Gly	Phe	Ala 240
Ile	Pro	Ile	Gly	Gln 245	Ala	Met	Ala	Ile	Ala 250	Gly	Gln	Ile	Arg	Ser 255	Gly
Gly	Gly	Ser	Pro 260	Thr	Val	His	Ile	Gly 265	Pro	Thr	Ala	Phe	Leu 270	Gly	Leu
Gly	Val	Val 275	Asp	Asn	Asn	Gly	Asn 280	Gly	Ala	Arg	Val	Gln 285	Arg	Val	Val
Gly	Ser 290	Ala	Pro	Ala	Ala	Ser 295	Leu	Gly	Ile	Ser	Thr 300		Asp	Val	Ile
Thr 305	Ala	Val	Asp	Gly	Ala 310	Pro	Ile	Asn	Ser	Ala 315	Thr	Ala	Met	Ala	320
Ala	Leu	Asn	Gly	His 325	His	Pro	Gly	Asp	Val 330	Ile	Ser	Val	Asn	Trp 335	
Thr	Lys	Ser	Gly 340	Gly	Thr	Arg	Thr	Gly 345	Asn	Val	Thr	Leu	Ala 350		Gl
Pro	Pro	Ala 355										•			

(2) INFORMATION FOR SEQ ID NO:81:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 205 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:81:

Ser Pro Lys Pro Asp Ala Glu Glu Gln Gly Val Pro Val Ser Pro Thr 1 5 10 15

Ala Ser Asp Pro Ala Leu Leu Ala Glu Ile Arg Gln Ser Leu Asp Ala 20 25 30

Thr Lys Gly Leu Thr Ser Val His Val Ala Val Arg Thr Thr Gly Lys 35 40 45

Val Asp Ser Leu Leu Gly Ile Thr Ser Ala Asp Val Asp Val Arg Ala 50 55 60

Asn Pro Leu Ala Ala Lys Gly Val Cys Thr Tyr Asn Asp Glu Gln Gly 65 70 75 80

Val Pro Phe Arg Val Gln Gly Asp Asn Ile Ser Val Lys Leu Phe Asp 85 90 95

Asp Trp Ser Asn Leu Gly Ser Ile Ser Glu Leu Ser Thr Ser Arg Val

Leu Asp Pro Ala Ala Gly Val Thr Gln Leu Leu Ser Gly Val Thr Asn 115 120 125

Leu Gln Ala Gln Gly Thr Glu Val Ile Asp Gly Ile Ser Thr Thr Lys 130 135 140

Ile Thr Gly Thr Ile Pro Ala Ser Ser Val Lys Met Leu Asp Pro Gly 145 150 155 160

Ala Lys Ser Ala Arg Pro Ala Thr Val Trp Ile Ala Gln Asp Gly Ser 165 170 175

His His Leu Val Arg Ala Ser Ile Asp Leu Gly Ser Gly Ser Ile Gln 180 185 190

Leu Thr Gln Ser Lys Trp Asn Glu Pro Val Asn Val Asp 195 200 205

(2) INFORMATION FOR SEQ ID NO:82:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 286 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:82:
- Gly Asp Ser Phe Trp Ala Ala Ala Asp Gln Met Ala Arg Gly Phe Val 1 5 10 15
- Leu Gly Ala Thr Ala Gly Arg Thr Thr Leu Thr Gly Glu Gly Leu Gln 20 25 30
- His Ala Asp Gly His Ser Leu Leu Leu Asp Ala Thr Asn Pro Ala Val 35 40 45
- Val Ala Tyr Asp Pro Ala Phe Ala Tyr Glu Ile Gly Tyr Ile Xaa Glu 50 55 60
- Ser Gly Leu Ala Arg Met Cys Gly Glu Asn Pro Glu Asn Ile Phe Phe 65 70 75 80
- Tyr Ile Thr Val Tyr Asn Glu Pro Tyr Val Gln Pro Pro Glu Pro Glu 85 90 95
- Asn Phe Asp Pro Glu Gly Val Leu Gly Gly Ile Tyr Arg Tyr His Ala 100 105 110
- Ala Thr Glu Gln Arg Thr Asn Lys Xaa Gln Ile Leu Ala Ser Gly Val 115 120 125
- Ala Met Pro Ala Ala Leu Arg Ala Ala Gln Met Leu Ala Ala Glu Trp 130 135 140
- Asp Val Ala Ala Asp Val Trp Ser Val Thr Ser Trp Gly Glu Leu Asn 145 150 155 160
- Arg Asp Gly Val Val Ile Glu Thr Glu Lys Leu Arg His Pro Asp Arg 165 170 175
- Pro Ala Gly Val Pro Tyr Val Thr Arg Ala Leu Glu Asn Ala Arg Gly 180 185 190
- Pro Val Ile Ala Val Ser Asp Trp Met Arg Ala Val Pro Glu Gln Ile 195 200 205
- Arg Pro Trp Val Pro Gly Thr Tyr Leu Thr Leu Gly Thr Asp Gly Phe 210 215 220
- Gly Phe Ser Asp Thr Arg Pro Ala Gly Arg Arg Tyr Phe Asn Thr Asp

225 230 235 240

Ala Glu Ser Gln Val Gly Arg Gly Phe Gly Arg Gly Trp Pro Gly Arg 245 250 255

Arg Val Asn Ile Asp Pro Phe Gly Ala Gly Arg Gly Pro Pro Ala Gln 260 265 270

Leu Pro Gly Phe Asp Glu Gly Gly Gly Leu Arg Pro Xaa Lys 275 280 285

(2) INFORMATION FOR SEQ ID NO:83:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 173 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:83:

Thr Lys Phe His Ala Leu Met Gln Glu Gln Ile His Asn Glu Phe Thr
1 5 10 15

Ala Ala Gln Gln Tyr Val Ala Ile Ala Val Tyr Phe Asp Ser Glu Asp 20 25 30

Leu Pro Gln Leu Ala Lys His Phe Tyr Ser Gln Ala Val Glu Glu Arg 35 40 45

Asn His Ala Met Met Leu Val Gln His Leu Leu Asp Arg Asp Leu Arg 50 55 60

Val Glu Ile Pro Gly Val Asp Thr Val Arg Asn Gln Phe Asp Arg Pro 65 70 75 80

Arg Glu Ala Leu Ala Leu Ala Leu Asp Gln Glu Arg Thr Val Thr Asp 85 90 95

Gln Val Gly Arg Leu Thr Ala Val Ala Arg Asp Glu Gly Asp Phe Leu 100 105 110

Gly Glu Gln Fhe Met Gln Trp Phe Leu Gln Glu Gln Ile Glu Glu Val 115 120 125

Ala Leu Met Ala Thr Leu Val Arg Val Ala Asp Arg Ala Gly Ala Asn 130 135 140

Leu Phe Glu Leu Glu Asn Phe Val Ala Arg Glu Val Asp Val Ala Pro 145 150 155 160

Ala Ala Ser Gly Ala Pro His Ala Ala Gly Gly Arg Leu 165 170

- (2) INFORMATION FOR SEQ ID NO:84:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 107 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:84:
 - Arg Ala Asp Glu Arg Lys Asn Thr Thr Met Lys Met Val Lys Ser Ile

 5 10 15
 - Ala Ala Gly Leu Thr Ala Ala Ala Ile Gly Ala Ala Ala Gly 20 25 30
 - Val Thr Ser Ile Met Ala Gly Gly Pro Val Val Tyr Gln Met Gln Pro 35 40 45
 - Val Val Phe Gly Ala Pro Leu Pro Leu Asp Pro Xaa Ser Ala Pro Xaa 50 55 60
 - Val Pro Thr Ala Ala Gln Trp Thr Xaa Leu Leu Asn Xaa Leu Xaa Asp 65 70 75 80
 - Pro Asn Val Ser Phe Xaa Asn Lys Gly Ser Leu Val Glu Gly Gly Ile 85 90 95
 - Gly Gly Xaa Glu Gly Xaa Xaa Arg Arg Xaa Gln 100 105
- (2) INFORMATION FOR SEQ ID NO:85:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 125 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:85:
 - Val Leu Ser Val Pro Val Gly Asp Gly Phe Trp Xaa Arg Val Val Asn
 1 10 15
 - Pro Leu Gly Gln Pro Ile Asp Gly Arg Gly Asp Val Asp Ser Asp Thr

20 25 30

Arg Arg Ala Leu Glu Leu Gln Ala Pro Ser Val Val Xaa Arg Gln Gly
35 40 45

Val Lys Glu Pro Leu Xaa Thr Gly Ile Lys Ala Ile Asp Ala Met Thr 50 55 60

Pro Ile Gly Arg Gly Gln Arg Gln Leu Ile Ile Gly Asp Arg Lys Thr 65 70 75 80

Gly Lys Asn Arg Arg Leu Cys Arg Thr Pro Ser Ser Asn Gln Arg Glu 85 90 95

Glu Leu Gly Val Arg Trp Ile Pro Arg Ser Arg Cys Ala Cys Val Tyr 100 105 110

Val Gly His Arg Ala Arg Arg Gly Thr Tyr His Arg Arg 115 120 125

(2) INFORMATION FOR SEQ ID NO:86:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 117 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:86:

Cys Asp Ala Val Met Gly Phe Leu Gly Gly Ala Gly Pro Leu Ala Val 1 5 10 15

Val Asp Gln Gln Leu Val Thr Arg Val Pro Gln Gly Trp Ser Phe Ala 20 25 30

Gln Ala Ala Val Pro Val Val Phe Leu Thr Ala Trp Tyr Gly Leu 35 40 45

Ala Asp Leu Ala Glu Ile Lys Ala Gly Glu Ser Val Leu Ile His Ala 50 55 60

Gly Thr Gly Gly Val Gly Met Ala Ala Val Gln Leu Ala Arg Gln Trp 65 70 75 80

Gly Val Glu Val Phe Val Thr Ala Ser Arg Gly Lys Trp Asp Thr Leu 85 90 95

Arg Ala Xaa Xaa Phe Asp Asp Xaa Pro Tyr Arg Xaa Phe Pro His Xaa 100 105 110

Arg Ser Ser Xaa Gly 115

- (2) INFORMATION FOR SEQ ID NO:87:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 103 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:87:

Met Tyr Arg Phe Ala Cys Arg Thr Leu Met Leu Ala Ala Cys Ile Leu 1 5 10 15

Ala Thr Gly Val Ala Gly Leu Gly Val Gly Ala Gln Ser Ala Ala Gln 20 25 30

Thr Ala Pro Val Pro Asp Tyr Tyr Trp Cys Pro Gly Gln Pro Phe Asp 35 40 45

Pro Ala Trp Gly Pro Asn Trp Asp Pro Tyr Thr Cys His Asp Asp Phe 50 55 60

His Arg Asp Ser Asp Gly Pro Asp His Ser Arg Asp Tyr Pro Gly Pro 65 70 75 80

Ile Leu Glu Gly Pro Val Leu Asp Asp Pro Gly Ala Ala Pro Pro Pro 85 90 95

Pro Ala Ala Gly Gly Gly Ala 100

- (2) INFORMATION FOR SEQ ID NO:88:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 88 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:88:

Val Gln Cys Arg Val Trp Leu Glu Ile Gln Trp Arg Gly Met Leu Gly
1 5 10 15

Ala Asp Gln Ala Arg Ala Gly Gly Pro Ala Arg Ile Trp Arg Glu His 20 25 30

Ser Met Ala Ala Met Lys Pro Arg Thr Gly Asp Gly Pro Leu Glu Ala 35 40 45

Thr Lys Glu Gly Arg Gly Ile Val Met Arg Val Pro Leu Glu Gly Gly 50 55 60

Gly Arg Leu Val Val Glu Leu Thr Pro Asp Glu Ala Ala Ala Leu Gly 65 70 75 80

Asp Glu Leu Lys Gly Val Thr Ser

(2) INFORMATION FOR SEQ ID NO:89:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 95 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:89:

Thr Asp Ala Ala Thr Leu Ala Gln Glu Ala Gly Asn Phe Glu Arg Ile 1 5 10 15

Ser Gly Asp Leu Lys Thr Gln Ile Asp Gln Val Glu Ser Thr Ala Gly

Ser Leu Gln Gly Gln Trp Arg Gly Ala Ala Gly Thr Ala Ala Gln Ala 35 40 45

Ala Val Val Arg Phe Gln Glu Ala Ala Asn Lys Gln Lys Gln Glu Leu 50 55 60

Asp Glu Ile Ser Thr Asn Ile Arg Gln Ala Gly Val Gln Tyr Ser Arg 65 70 75 80

Ala Asp Glu Glu Gln Gln Gln Ala Leu Ser Ser Gln Met Gly Phe 85 90 95

(2) INFORMATION FOR SEQ ID NO:90:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 166 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:90:

Arg Ala Asn Glu Val Glu Ala Pro Met Ala Asp Pro Pro Thr Asp Val 20 25 30

Pro Ile Thr Pro Cys Glu Leu Thr Xaa Xaa Lys Asn Ala Ala Gln Gln 35 40 45

Xaa Val Leu Ser Ala Asp Asn Met Arg Glu Tyr Leu Ala Ala Gly Ala 50 55 60

Lys Glu Arg Gln Arg Leu Ala Thr Ser Leu Arg Asn Ala Ala Lys Xaa 65 70 75 80

Tyr Gly Glu Val Asp Glu Glu Ala Ala Thr Ala Leu Asp Asn Asp Gly 85 90 95

Glu Gly Thr Val Gln Ala Glu Ser Ala Gly Ala Val Gly Gly Asp Ser

Ser Ala Glu Leu Thr Asp Thr Pro Arg Val Ala Thr Ala Gly Glu Pro 115 120 125

Asn Phe Met Asp Leu Lys Glu Ala Ala Arg Lys Leu Glu Thr Gly Asp 130 135 140

Gln Gly Ala Ser Leu Ala His Xaa Gly Asp Gly Trp Asn Thr Xaa Thr 145 150 155 160

Leu Thr Leu Gln Gly Asp

(2) INFORMATION FOR SEQ ID NO:91:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 5 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:91:

Arg Ala Glu Arg Met

1

5

(2) INFORMATION FOR SEQ ID NO:92:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 263 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 92:

Val Ala Trp Met Ser Val Thr Ala Gly Gln Ala Glu Leu Thr Ala Ala 1 5 10 15/

Gln Val Arg Val Ala Ala Ala Ala Tyr Glu Thr Ala Tyr Gly Leu Thr 20 25 30

Val Pro Pro Pro Val Ile Ala Glu Asn Arg Ala Glu Leu Met Ile Leu 35 40 45

Ile Ala Thr Asn Leu Leu Gly Gln Asn Thr Pro Ala Ile Ala Val Asn 50 55 60

Glu Ala Glu Tyr Gly Glu Met Trp Ala Gln Asp Ala Ala Ala Met Phe
65 70 75 80

Gly Tyr Ala Ala Ala Thr Ala Thr Ala Thr Ala Thr Leu Leu Pro Phe 85 90 95

Glu Glu Ala Pro Glu Met Thr Ser Ala Gly Gly Leu Leu Glu Gln Ala 100 105 110

Ala Ala Val Glu Glu Ala Ser Asp Thr Ala Ala Ala Asn Gln Leu Met 115 120 125

Asn Asn Val Pro Gln Ala Leu Lys Gln Leu Ala Gln Pro Thr Gln Gly 130 135 140

Thr Thr Pro Ser Ser Lys Leu Gly Gly Leu Trp Lys Thr Val Ser Pro 145 150 155 160

His Arg Ser Pro Ile Ser Asn Met Val Ser Met Ala Asn Asn His Met 165 170 . 175

Ser Met Thr Asn Ser Gly Val Ser Met Thr Asn Thr Leu Ser Ser Met 180 185 190

Leu Lys Gly Phe Ala Pro Ala Ala Ala Ala Gln Ala Val Gln Thr Ala 195 200 205

Ala Gln Asn Gly Val Arg Ala Met Ser Ser Leu Gly Ser Ser Leu Gly

210 215 220

Ser Ser Gly Leu Gly Gly Gly Val Ala Ala Asn Leu Gly Arg Ala Ala 225 230 235 240

Ser Val Arg Tyr Gly His Arg Asp Gly Gly Lys Tyr Ala Xaa Ser Gly 245 250 255

Arg Arg Asn Gly Gly Pro Ala 260

(2) INFORMATION FOR SEQ ID NO:93:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 303 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:93:

Met Thr Tyr Ser Pro Gly Asn Pro Gly Tyr Pro Gln Ala Gln Pro Ala 1 5 10 15

Gly Ser Tyr Gly Gly Val Thr Pro Ser Phe Ala His Ala Asp Glu Gly 20 25 30

Ala Ser Lys Leu Pro Met Tyr Leu Asn Ile Ala Val Ala Val Leu Gly 35 40 45

Leu Ala Ala Tyr Phe Ala Ser Phe Gly Pro Met Phe Thr Leu Ser Thr 50 55 60

Glu Leu Gly Gly Gly Asp Gly Ala Val Ser Gly Asp Thr Gly Leu Pro
65 70 . 75 80

Val Gly Val Ala Leu Leu Ala Ala Leu Leu Ala Gly Val Val Leu Val 85 90 95

Pro Lys Ala Lys Ser His Val Thr Val Val Ala Val Leu Gly Val Leu 100 105 110

Gly Val Phe Leu Met Val Ser Ala Thr Phe Asn Lys Pro Ser Ala Tyr 115 120 125

Ser Thr Gly Trp Ala Leu Trp Val Val Leu Ala Phe Ile Val Phe Gln 130 135 140

Ala Val Ala Ala Val Leu Ala Leu Leu Val Glu Thr Gly Ala Ile Thr 145 150 155 160

Ala	Pro	Ala	Pro	Arg 165	Pro	Lys	Phe	Asp	Pro 170	Tyr	Gly	Gln	Tyr	Gly 175	Arg
Tyr	Gly	Gln	Tyr 180	Gly	Gln	Tyr	Gly	Val 185	Gln	Pro	Gly	Gly	Tyr 190	Tyr	Gly
Gln	Gln	Gly 195	Ala	Gln	Gln	Ala	Ala 200	Gly	Leu	Gln	Ser	Pro 205	Gly	Pro	Glr
Gln	Ser 210	Pro	Gln	Pro	Pro	Gly 215	Tyr	Gly	Ser	Gln	Tyr 220	Gly	Gly	Tyr	Ser
Ser 225	Ser	Pro	Ser	Gln	Ser 230	Gly	Ser	Gly	Tyr	Thr 235	Ala	Gln	Pro	Pro	Ala 240
Gln	Pro	Pro	Ala	Gln 245	Ser	Gly	Ser	Gln	Gln 250	Ser	His	Gln	Gly	Pro 255	
Thr	Pro	Pro	Thr 260	Gly	Phe	Pro	Ser	Phe 265	Ser	Pro	Pro	Pro	Pro 270	Val ^{//}	Ser
Ala	Gly	Thr 275	Gly	Ser	Gln	Ala	Gly 280	Ser	Ala	Pro	Val	Asn 285	Tyr	Ser	Asr
Pro	Ser 290	Gly	Gly	Glu	Gln	Ser 295	Ser	Ser	Pro	Gly	Gly 300	Ala	Pro	Val	

(2) INFORMATION FOR SEQ ID NO:94:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 507 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:94:

ATGAAGATGG	TGAAATCGAT	CGCCGCAGGT	CTGACCGCCG	CGGCTGCAAT	CGGCGCCGCT	60
GCGGCCGGTG	TGACTTCGAT	CATGGCTGGC	GGCCCGGTCG	TATACCAGAT	GCAGCCGGTC	120
GTCTTCGGCG	CGCCACTGCC	GTTGGACCCG	GCATCCGCCC	CTGACGTCCC	GACCGCCGCC	180
CAGTTGACCA	GCCTGCTCAA	CAGCCTCGCC	GATCCCAACG	TGTCGTTTGC	GAACAAGGGC	240
AGTCTGGTCG	AGGGCGGCAT	CGGGGCACC	GAGGCGCGCA	TCGCCGACCA	CAAGCTGAAG	300
AAGGCCGCCG	AGCACGGGGA	TCTGCCGCTG	TCGTTCAGCG	TGACGAACAT	CCAGCCGGCG	360
GCCGCCGGTT	CGGCCACCGC	CGACGTTTCC	GTCTCGGGTC	CGAAGCTCTC	GTCGCCGGTC	420

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128

ACGCAGAACG TCACGTTCGT GAATCAAGGC GGCTGGATGC TGTCACGCGC ATCGGCGATG 480
GAGTTGCTGC AGGCCGCAGG GAACTGA 507

(2) INFORMATION FOR SEQ ID NO:95:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 168 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:95:

Met Lys Met Val Lys Ser Ile Ala Ala Gly Leu Thr Ala Ala Ala Ala 1 5 10 15

Ile Gly Ala Ala Ala Gly Val Thr Ser Ile Met Ala Gly Gly Pro $20 \hspace{1.5cm} 25 \hspace{1.5cm} 30$

Val Val Tyr Gln Met Gln Pro Val Val Phe Gly Ala Pro Leu Pro Leu 35 40 45

Asp Pro Ala Ser Ala Pro Asp Val Pro Thr Ala Ala Gln Leu Thr Ser 50 55 60

Leu Leu Asn Ser Leu Ala Asp Pro Asn Val Ser Phe Ala Asn Lys Gly 65 70 75 80

Ser Leu Val Glu Gly Gly Ile Gly Gly Thr Glu Ala Arg Ile Ala Asp 85 90 95

His Lys Leu Lys Lys Ala Ala Glu His Gly Asp Leu Pro Leu Ser Phe 100 105 110

Ser Val Thr Asn Ile Gln Pro Ala Ala Gly Ser Ala Thr Ala Asp 115 120 125

Val Ser Val Ser Gly Pro Lys Leu Ser Ser Pro Val Thr Gln Asn Val 130 135 140

Thr Phe Val Asn Gln Gly Gly Trp Met Leu Ser Arg Ala Ser Ala Met 145 150 155 160

Glu Leu Leu Gln Ala Ala Gly Asn 165

(2) INFORMATION FOR SEQ ID NO:96:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 500 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:96:

CGTGGCAATG	TCGTTGACCG	TCGGGGCCGG	GGTCGCCTCC	GCAGATCCCG	TGGACGCGGT	60
CATTAACACC	ACCTGCAATT	ACGGGCAGGT	AGTAGCTGCG	CTCAACGCGA	CGGATCCGGG	120
GGCTGCCGCA	CAGTTCAACG	CCTCACCGGT	GGCGCAGTCC	TATTTGCGCA	ATTTCCTCGC	180
CGCACCGCCA	CCTCAGCGCG	CTGCCATGGC	CGCGCAATTG	CAAGCTGTGC	CGGGGGCGC	240
ACAGTACATC	GGCCTTGTCG	AGTCGGTTGC	CGGCTCCTGC	AACAACTATT	AAGCCCATGC	300
GGGCCCCATC	CCGCGACCCG	GCATCGTCGC	CGGGGCTAGG	CCAGATTGCC	CCGCTCCTCA	360
ACGGGCCGCA	TCCCGCGACC	CGGCATCGTC	GCCGGGGCTA	GGCCAGATTG	CCCCGCTCCT	420
CAACGGGCCG	CATCTCGTGC	CGAATTCCTG	CAGCCCGGGG	GATCCACTAG	TTCTAGAGCG	480
GCCGCCACCG	CGGTGGAGCT					500

(2) INFORMATION FOR SEQ ID NO:97:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 96 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:97:

Val Ala Met Ser Leu Thr Val Gly Ala Gly Val Ala Ser Ala Asp Pro 1 5 10 15

Val Asp Ala Val Ile Asn Thr Thr Cys Asn Tyr Gly Gln Val Val Ala 20 25 30

Ala Leu Asn Ala Thr Asp Pro Gly Ala Ala Ala Gln Phe Asn Ala Ser 35 40 45

Pro Val Ala Gln Ser Tyr Leu Arg Asn Phe Leu Ala Ala Pro Pro Pro 50 55 60

130

Gln Arg Ala Ala Met Ala Ala Gln Leu Gln Ala Val Pro Gly Ala A	GIN	in Arg Ala	J Ala Ala Met Ala	Ala Gln Leu	Gln Ala Val	Pro Gly	777	
	65		70		25	IIO CIY	ura	мта
75			. •		75			80

Gln Tyr Ile Gly Leu Val Glu Ser Val Ala Gly Ser Cys Asn Asn Tyr

- (2) INFORMATION FOR SEQ ID NO:98:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 154 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:98: ATGACAGAGC AGCAGTGGAA TTTCGCGGGT ATCGAGGCCG CGGCAAGCGC AATCCAGGGA . 60 AATGTCACGT CCATTCATTC CCTCCTTGAC GAGGGGAAGC AGTCCCTGAC CAAGCTCGCA 120 GCGGCCTGGG GCGGTAGCGG TTCGGAAGCG TACC
- (2) INFORMATION FOR SEQ ID NO:99:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 51 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:99:
 - Met Thr Glu Gln Gln Trp Asn Phe Ala Gly Ile Glu Ala Ala Ser
 - Ala Ile Gln Gly Asn Val Thr Ser Ile His Ser Leu Leu Asp Glu Gly 25
 - Lys Gln Ser Leu Thr Lys Leu Ala Ala Trp Gly Gly Ser Gly Ser

Glu Ala Tyr 50

- (2) INFORMATION FOR SEQ ID NO:100:
 - (i) SEQUENCE CHARACTERISTICS:

(A)	LENGTH:	282 ba	ase	pairs
(B)	TYPE: nu	ucleic	aci	id
	CODANDE	MECC.		1-

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:100:

CGGTCGCGCA CTTCCAGGTG ACTATGAAAG TCGGCTTCCG NCTGGAGGAT TCCTGAACCT 60

TCAAGCGCGG CCGATAACTG AGGTGCATCA TTAAGCGACT TTTCCAGAAC ATCCTGACGC 120

GCTCGAAACG CGGCACAGCC GACGGTGGCT CCGNCGAGGC GCTGNCTCCA AAATCCCTGA 180

GACAATTCGN CGGGGGCGCC TACAAGGAAG TCGGTGCTGA ATTCGNCGNG TATCTGGTCG 240

ACCTGTGTGG TCTGNAGCCG GACGAAGCGG TGCTCGACGT CG 282

(2) INFORMATION FOR SEQ ID NO:101:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 3058 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:101:

GATCGTACCC GTGCGAGTGC TCGGGCCGTT TGAGGATGGA GTGCACGTGT CTTTCGTGAT 60 GGCATACCCA GAGATGTTGG CGGCGGCGGC TGACACCCTG CAGAGCATCG GTGCTACCAC 120 TGTGGCTAGC AATGCCGCTG CGGCGGCCCC GACGACTGGG GTGGTGCCCC CCGCTGCCGA 180 TGAGGTGTCG GCGCTGACTG CGGCGCACTT CGCCGCACAT GCGGCGATGT ATCAGTCCGT 240 GAGCGCTCGG GCTGCTGCGA TTCATGACCA GTTCGTGGCC ACCCTTGCCA GCAGCGCCAG 300 CTCGTATGCG GCCACTGAAG TCGCCAATGC GGCGGCGGCC AGCTAAGCCA GGAACAGTCG 360 GCACGAGAAA CCACGAGAAA TAGGGACACG TAATGGTGGA TTTCGGGGCG TTACCACCGG 420 AGATCAACTC CGCGAGGATG TACGCCGGCC CGGGTTCGGC CTCGCTGGTG GCCGCGGCTC 480 AGATGTGGGA CAGCGTGGCG AGTGACCTGT TTTCGGCCGC GTCGGCGTTT CAGTCGGTGG 540 TCTGGGGTCT GACGGTGGGG TCGTGGATAG GTTCGTCGGC GGGTCTGATG GTGGCGGCGG 600

CCTCGCCGTA	TGTGGCGTGG	ATGAGCGTCA	CCGCGGGGCA	GGCCGAGCTG	ACCGCCGCCC	660
AGGTCCGGGT	TGCTGCGGCG	GCCTACGAGA	CGGCGTATGG	GCTGACGGTG	cccccccc	720
TGATCGCCGA	GAACCGTGCT	GAACTGATGA	TTCTGATAGC	GACCAACCTC	TTGGGGCAAA	780
ACACCCCGGC	GATCGCGGTC	AACGAGGCCG	AATACGGCGA	GATGTGGGCC	CAAGACGCCG	840
CCGCGATGTT	TGGCTACGCC	GCGGCGACGG	CGACGGCGAC	GGCGACGTTG	CTGCCGTTCG	900
AGGAGGCGCC	GGAGATGACC	AGCGCGGGTG	GGCTCCTCGA	GCAGGCCGCC	GCGGTCGAGG	960
AGGCCTCCGA	CACCGCCGCG	GCGAACCAGT	TGATGAACAA	TGTGCCCCAG	GCGCTGCAAC	1020
AGCTGGCCCA	GCCCACGCAG	GGCACCACGC	CTTCTTCCAA	GCTGGGTGGC	CTGTGGAAGA	1080
CGGTCTCGCC	GCATCGGTCG	CCGATCAGCA	ACATGGTGTC	GATGGCCAAC	AACCACATGT	1140
CGATGACCAA	CTCGGGTGTG	TCGATGACCA	ACACCTTGAG	CTCGATGTTG	AAGGGCTTTG	1200
CTCCGGCGGC	GGCCGCCCAG	GCCGTGCAAA	CCGCGGCGCA	AAACGGGGTC	CGGGCGATGA	1260
GCTCGCTGGG	CAGCTCGCTG	GGTTCTTCGG	GTCTGGGCGG	TGGGGTGGCC	GCCAACTTGG	1320
GTCGGGCGGC	CTCGGTCGGT	TCGTTGTCGG	TGCCGCAGGC	CTGGGCCGCG	GCCAACCAGG	1380
CAGTCACCCC	GGCGGCGCGG	GCGCTGCCGC	TGACCAGCCT	GACCAGCGCC	GCGGAAAGAG	1440
GGCCCGGGCA	GATGCTGGGC	GGGCTGCCGG	TGGGGCAGAT	GGGCGCCAGG	GCCGGTGGTG	1500
GGCTCAGTGG	TGTGCTGCGT	GTTCCGCCGC	GACCCTATGT	GATGCCGCAT	TCTCCGGCGG	1560
CCGGCTAGGA	GAGGGGGCGC	AGACTGTCGT	TATTTGACCA	GTGATCGGCG	GTCTCGGTGT	1620
TTCCGCGGCC	GGCTATGACA	ACAGTCAATG	TGCATGACAA	GTTACAGGTA	TTAGGTCCAG	1680
GTTCAACAAG	GAGACAGGCA	ACATGGCCTC	ACGTTTTATG	ACGGATCCGC	ACGCGATGCG	1740
GGACATGGCG	GGCCGTTTTG	AGGTGCACGC	CCAGACGGTG	GAGGACGAGG	CTCGCCGGAT	1800
GTGGGCGTCC	GCGCAAAACA	TTTCCGGTGC	GGGCTGGAGT	GGCATGGCCG	AGGCGACCTC	1860
GCTAGACACC	ATGGCCCAGA	TGAATCAGGC	GTTTCGCAAC	ATCGTGAACA	TGCTGCACGG	1920
GGTGCGTGAC	GGGCTGGTTC	GCGACGCCAA	CAACTACGAG	CAGCAAGAGC	AGGCCTCCCA	1980
GCAGATCCTC	AGCAGCTAAC	GTCAGCCGCT	GCAGCACAAT	ACTTTTACAP	GCGAAGGAGA	2040
ACAGGTTCGA	TGACCATCAA	CTATCAATTC	GGGGATGTCG	ACGCTCACGG	CGCCATGATC	2100
CGCGCTCAGG	CCGGGTTGCT	GGAGGCCGAG	CATCAGGCCA	TCATTCGTGA	TGTGTTGACC	2160
GCGAGTGACT	TTTGGGGCGG	CGCCGGTTCG	GCGGCCTGCC	AGGGGTTCAT	TACCCAGTTG	2220
GGCCGTAACT	тссасстсат	CTACGAGCAG	GCCAACGCCC	: ACGGGCAGAF	GGTGCAGGCT	2280

GCCGGCAACA	ACATGGCGCA	AACCGACAGC	GCCGTCGGCT	CCAGCTGGGC	CTGACACCAG	2340
GCCAAGGCCA	GGGACGTGGT	GTACGAGTGA	AGTTCCTCGC	GTGATCCTTC	GGGTGGCAGT	2400
CTAAGTGGTC	AGTGCTGGGG	TGTTGGTGGT	TTGCTGCTTG	GCGGGTTCTT	CGGTGCTGGT	2460
CAGTGCTGCT	CGGGCTCGGG	TGAGGACCTC	GAGGCCCAGG	TAGCGCCGTC	CTTCGATCCA	2520
TTCGTCGTGT	TGTTCGGCGA	GGACGGCTCC	GACGAGGCGG	ATGATCGAGG	CGCGGTCGGG	2580
GAAGATGCCC	ACGACGTCGG	TTCGGCGTCG	TACCTCTCGG	TTGAGGCGTT	CCTGGGGGTT	2640
GTTGGACCAG	ATTTGGCGCC	AGATCTGCTT	GGGGAAGGCG	GTGAACGCCA	GCAGGTCGGT	2700
GCGGGCGGTG	TCGAGGTGCT	CGGCCACCGC	GGGGAGTTTG	TCGGTCAGAG	CGTCGAGTAC	2760
CCGATCATAT	TGGGCAACAA	CTGATTCGGC	GTCGGGCTGG	TCGTAGATGG	AGTGCAGCÁG	2820
GGTGCGCACC	CACGGCCAGG	AGGGCTTCGG	GGTGGCTGCC	ATCAGATTGG	CTGCGTAGTG	2880
GGTTCTGCAG	CGCTGCCAGG	CCGCTGCGGG	CAGGGTGGCG	CCGATCGCGG	CCACCAGGCC	2940
GGCGTGGGCG	TCGCTGGTGA	CCAGCGCGAC	CCCGGACAGG	CCGCGGGCGA	CCAGGTCGCG	3000
GAAGAACGCC	AGCCAGCCGG	CCCCGTCCTC	GGCGGAGGTG	ACCTGGATGC	CCAGGATC	3058

(2) INFORMATION FOR SEQ ID NO:102:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 391 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:102:

Met Val Asp Phe Gly Ala Leu Pro Pro Glu Ile Asn Ser Ala Arg Met 1 5 10 15

Tyr Ala Gly Pro Gly Ser Ala Ser Leu Val Ala Ala Ala Gln Met Trp 20 25 30

Asp Ser Val Ala Ser Asp Leu Phe Ser Ala Ala Ser Ala Phe Gln Ser 35 40 45

Val Val Trp Gly Leu Thr Val Gly Ser Trp Ile Gly Ser Ser Ala Gly 50 55 60

Leu Met Val Ala Ala Ala Ser Pro Tyr Val Ala Trp Met Ser Val Thr 65 70 75 80

A	la	G1	y G	ln i	Ala	G1 85	u Le	u T	hr	Ala	a Al	.a G	Sln 90	Va.	l Ar	g Va	al Al	la A 9		Ala
A.	la	Ty:	r G	lu :	hr 100	Ala	а Ту	r G	ly	Let	1 Th	r V 5	'al	Pro	o Pr	o Pr	0 Va		le	Ala
G]	lu	Ası	n Ai 11	rg <i>I</i> .5	lla	Glı	ı Le	u Me	et	11e 120	e Le	u I	le	Ala	a Th	r As 12	n Le	u Le	eu	Gly
G1	n.	Asr 130	n Th	r F	ro	Ala	ı Il	e Al	La 35	Val	Ası	n G	lu	Ala	Gl: 140	а Ту О	r Gl	y Gl	.u	Met
13	J						120	J						155	•		a Al			160
						103						1	70				o Gl	1/7	5	
				1	30						185	Ò			•		u Gl: 19	0		
			19.	,					2	200						20!				
	2	10						21.	5						220		Se:			
							230							235) Ile		2	240
					4	245						25	0				Ser	25	5	٠
				26	U						265						Ala 270)		
		•	215						28	80						285				
	د د	, 0						293							300		Gly			
303						•	310						3	315			Leu		3	20
					3	25						330)				Ala	335		-
				340						3	345						Gly 350			_
Gln	Me	t L 3	eu 55	Gly	G1	ly I	eu :	Pro	Va 36	1 G 0	Зly	Gln	М	et (Ala 365	Arg	Ala	G.	ly

Gly Gly Leu Ser Gly Val Leu Arg Val Pro Pro Arg Pro Tyr Val Met 370 375 380

Pro His Ser Pro Ala Ala Gly 385 390

(2) INFORMATION FOR SEQ ID NO:103:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1725 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:103:

GACGTCAGCA CCCGCCGTGC AGGGCTGGAG CGTGGTCGGT TTTGATCTGC GGTCAAGGTG 60 ACGTCCCTCG GCGTGTCGCC GGCGTGGATG CAGACTCGAT GCCGCTCTTT AGTGCAACTA 120 ATTTCGTTGA AGTGCCTGCG AGGTATAGGA CTTCACGATT GGTTAATGTA GCGTTCACCC 180 CGTGTTGGGG TCGATTTGGC CGGACCAGTC GTCACCAACG CTTGGCGTGC GCGCCAGGCG 240 GGCGATCAGA TCGCTTGACT ACCAATCAAT CTTGAGCTCC CGGGCCGATG CTCGGGCTAA 300 ATGAGGAGGA GCACGCGTGT CTTTCACTGC GCAACCGGAG ATGTTGGCGG CCGCGGCTGG 360 CGAACTTCGT TCCCTGGGGG CAACGCTGAA GGCTAGCAAT GCCGCCGCAG CCGTGCCGAC 420 GACTGGGGTG GTGCCCCGG CTGCCGACGA GGTGTCGCTG CTGCTTGCCA CACAATTCCG 480 TACGCATGCG GCGACGTATC AGACGGCCAG CGCCAAGGCC GCGGTGATCC ATGAGCAGTT 540 TGTGACCACG CTGGCCACCA GCGCTAGTTC ATATGCGGAC ACCGAGGCCG CCAACGCTGT 600 GGTCACCGGC TAGCTGACCT GACGGTATTC GAGCGGAAGG ATTATCGAAG TGGTGGATTT 660 CGGGGCGTTA CCACCGGAGA TCAACTCCGC GAGGATGTAC GCCGGCCCGG GTTCGGCCTC 720 GCTGGTGGCC GCCGCGAAGA TGTGGGACAG CGTGGCGAGT GACCTGTTTT CGGCCGCGTC 780 GGCGTTTCAG TCGGTGGTCT GGGGTCTGAC GGTGGGGTCG TGGATAGGTT CGTCGGCGGG 840 TCTGATGGCG GCGGCGCCT CGCCGTATGT GGCGTGGATG AGCGTCACCG CGGGGCAGGC 900 CCAGCTGACC GCCGCCCAGG TCCGGGTTGC TGCGGCGGCC TACGAGACAG CGTATAGGCT 960 GACGGTGCCC CCGCCGGTGA TCGCCGAGAA CCGTACCGAA CTGATGACGC TGACCGCGAC 1020 CAACCTCTTG GGGCAAAACA CGCCGGCGAT CGAGGCCAAT CAGGCCGCAT ACAGCCAGAT 1080

GTGGGGCCA	A GACGCGGAGG	G CGATGTATGO	G CTACGCCGCC	ACGGCGGCGA	CGGCGACCGA	1140
GGCGTTGCT	G CCGTTCGAGO	ACGCCCCACT	GATCACCAAC	CCCGGCGGGC	TCCTTGAGCA	1200
GGCCGTCGC	G GTCGAGGAGG	CCATCGACAC	CGCCGCGGCG	AACCAGTTGA	TGAACAATGT	1260
GCCCCAAGCC	G CTGCAACAGC	TGGCCCAGCC	AGCGCAGGGC	GTCGTACCTT	CTTCCAAGCT	1320
GGGTGGGCTG	TGGACGGCGG	TCTCGCCGCA	TCTGTCGCCG	CTCAGCAACG	TCAGTTCGAT	1380
AGCCAACAAC	CACATGTCGA	TGATGGGCAC	GGGTGTGTCG	ATGACCAACA	CCTTGCACTC	1,440
GATGTTGAAG	GGCTTAGCTC	CGGCGGCGGC	TCAGGCCGTG	GAAACCGCGG	CGGAAAACGG	1500
GGTCTGGGCG	ATGAGCTCGC	TGGGCAGCCA	GCTGGGTTCG	TCGCTGGGTT	CTTCGGGTCT	1560
GGGCGCTGGG	GTGGCCGCCA	ACTTGGGTCG	GGCGGCCTCG	GTCGGTTCGT	TGTCGGTGCC	1620
GCCAGCATGG	GCCGCGGCCA	ACCAGGCGGT	CACCCGGCG	GCGCGGGCGC	TGCCGCTGAC	1680
CAGCCTGACC	AGCGCCGCCC	AAACCGCCCC	CGGACACATG	CTGGG		1725
/2\ TNT000						

(2) INFORMATION FOR SEQ ID NO:104:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 359 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS:
- (D) TOPCLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:104:

Val Val Asp Phe Gly Ala Leu Pro Pro Glu Ile Asn Ser Ala Arg Met 1 5 10 15

Tyr Ala Gly Pro Gly Ser Ala Ser Leu Val Ala Ala Ala Lys Met Trp 20 25 30

Asp Ser Val Ala Ser Asp Leu Phe Ser Ala Ala Ser Ala Phe Gln Ser 35 40 45

Val Val Trp Gly Leu Thr Val Gly Ser Trp Ile Gly Ser Ser Ala Gly 50 55 60

Leu Met Ala Ala Ala Ser Pro Tyr Val Ala Trp Met Ser Val Thr 65 70 75 80

Ala Gly Gln Ala Gln Leu Thr Ala Ala Gln Val Arg Val Ala Ala Ala 85 90 95

Ala	Туг	Glu	Thr 100	Ala	Tyr	Arg	Leu	Thr 105	Val	Pro	Pro	Pro	Val 110	Ile	Ala
Glu	Asn	Arg 115	Thr	Glu	Leu	Met	Thr 120	Leu	Thr	Ala	Thr	Asn 125	Leu	Leu	Gly
Gln	Asn 130	Thr	Pro	Ala	Ile	Glu 135	Ala	Asn	Gln	Ala	Ala 140	Tyr	Ser	Gln	Met
Trp 145	Gly	Gln	Asp	Ala	Glu 150	Ala	Met	Tyr	Gly	Tyr 155	Ala	Ala	Thr	Ala	Ala 160
Thr	Ala	Thr	Glu	Ala 165	Leu	Leu	Pro	Phe	Glu 170	Asp	Ala	Pro	Leu	Ile 175	Thr
Asn	Pro	Gly	Gly 180	Leu	Leu	Glu	Gln	Ala 185	Val	Ala	Val	Glu	Glu 190	Ala	Ile
Asp	Thr	Ala 195	Ala	Ala	Asn	Gln	Leu 200	Met	Asn	Asn	Val	Pro 205	Gln	Alá	Leu
Gln	Gln 210	Leu	Ala	Gln	Pro	Ala 215	Gln	Gly	Val	Val	Pro 220	Ser	Ser	Lys	Leu
Gly 225	Gly	Leu	Trp	Thr	Ala 230	Val	Ser	Pro	His	Leu 235	Ser	Pro	Leu	Ser	Asn 240
Val	Ser	Ser	Ile	Ala 245	Asn	Asn	His	Met	Ser 250	Met	Met	Gly	Thr	Gly 255	Val
Ser	Met	Thr	Asn 260	Thr	Leu	His	Ser	Met 265	Leu	Lys	Gly	Leu	Ala 270	Pro	Ala
Ala	Ala	Gln 275	Ala	Val	Glu	Thr	Ala 280	Ala	Glu	Asn	Gly	Val 285	Trp	Ala	Met
Ser	Ser 290	Leu	Gly	Ser	Gln	Leu 295	Gly	Ser	Ser	Leu	Gly 300	Ser	Ser	Gly	Leu
Gly 305	Ala	Gly	Val	Ala	Ala 310	Asn	Leu	Gly	Arg	Ala 315	Ala	Ser	Val	Gly	Ser 320
Leu	Ser	Val	Pro	Pro 325	Ala	Trp	Ala	Ala	Ala 330	Asn	Gln	Ala	Val	Thr 335	Pro
Ala	Ala	Arg	Ala 340	Leu	Pro	Leu	Thr	Ser 345	Leu	Thr	Ser	Ala	Ala 350	Gln	Thr
Ala	Pro	Gly 355	His	Met	Leu	Gly									

- (2) INFORMATION FOR SEQ ID NO:105:
 - (i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 3027 base pairs

- (B) TYPE: nucleic acid(C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:105:

AGTTCAGTCG AGAATGATAC TGACGGGCTG TATCCACGAT GGCTGAGACA ACCGAACCAC	60
CGTCGGACGC GGGGACATCG CAAGCCGACG CGATGGCGTT GGCCGCCGAA GCCGAAGCCG	120
CCGAAGCCGA AGCGCTGGCC GCCGCGCGC GGGCCCGTGC CCGTGCCGCC CGGTTGAAGC	180
GTGAGGCGCT GGCGATGGCC CCAGCCGAGG ACGAGAACGT CCCCGAGGAT ATGCAGACTG	240
GGAAGACGCC GAAGACTATG ACGACTATGA CGACTATGAG GCCGCAGACC AGGAGGCCGC	300
ACGGTCGGCA TCCTGGCGAC GGCGGTTGCG GGTGCGGTTA CCAAGACTGT CCACGATTGC	360
CATGGCGGCC GCAGTCGTCA TCATCTGCGG CTTCACCGGG CTCAGCGGAT ACATTGTGTG	420
GCAACACCAT GAGGCCACCG AACGCCAGCA GCGCGCCGCG GCGTTCGCCG CCGGAGCCAA	480
GCAAGGTGTC ATCAACATGA CCTCGCTGGA CTTCAACAAG GCCAAAGAAG ACGTCGCGCG	540
TGTGATCGAC AGCTCCACCG GCGAATTCAG GGATGACTTC CAGCAGCGGG CAGCCGATTT	600
CACCAAGGTT GTCGAACAGT CCAAAGTGGT CACCGAAGGC ACGGTGAACG CGACAGCCGT	660
CGAATCCATG AACGAGCATT CCGCCGTGGT GCTCGTCGCG GCGACTTCAC GGGTCACCAA	720
TTCCGCTGGG GCGAAAGACG AACCACGTGC GTGGCGGCTC AAAGTGACCG TGACCGAAGA	780
GGGGGGACAG TACAAGATGT CGAAAGTTGA GTTCGTACCG TGACCGATGA CGTACGCGAC	840
GTCAACACCG AAACCACTGA CGCCACCGAA GTCGCTGAGA TCGACTCAGC CGCAGGCGAA	900
GCCGGTGATT CGGCGACCGA GGCATTTGAC ACCGACTCTG CAACGGAATC TACCGCGCAG	960
AAGGGTCAGC GGCACCGTGA CCTGTGGCGA ATGCAGGTTA CCTTGAAACC CGTTCCGGTG	1020
ATTCTCATCC TGCTCATGTT GATCTCTGGG GGCGCGACGG GATGGCTATA CCTTGAGCAA	1080
TACGACCCGA TCAGCAGACG GACTCCGGCG CCGCCCGTGC TGCCGTCGCC GCGGCGTCTG	1140
ACGGGACAAT CGCGCTGTTG TGTATTCACC CGACACGTCG ACCAAGACTT CGCTACCGCC	1200
AGGTCGCACC TCGCCGGCGA TTTCCTGTCC TATACGACCA GTTCACGCAG CAGATCGTGG	1260
CTCCGGCGGC CAAACAGAAG TCACTGAAAA CCACCGCCAA GGTGGTGCGC GCGGCCGTGT	1320
CGGAGCTACA TCCGGATTCG GCCGTCGTTC TGGTTTTTGT CGACCAGAGC ACTACCAGTA	1380

AGGACAGCCC	CAATCCGTCG	ATGGCGGCCA	GCAGCGTGAT	GGTGACCCTA	GCCAAGGTCG	1440
ACGGCAATTG	GCTGATCACC	AAGTTCACCC	CGGTTTAGGT	TGCCGTAGGC	GGTCGCCAAG	1500
TCTGACGGGG	GCGCGGGTGG	CTGCTCGTGC	GAGATACCGG	CCGTTCTCCG	GACAATCACG	1560
GCCCGACCTC	AAACAGATCT	CGGCCGCTGT	CTAATCGGCC	GGGTTATTTA	AGATTAGTTG	1620
CCACTGTATT	TACCTGATGT	TCAGATTGTT	CAGCTGGATT	TAGCTTCGCG	GCAGGGCGGC	1680
TGGTGCACTT	TGCATCTGGG	GTTGTGACTA	CTTGAGAGAA	TTTGACCTGT	TGCCGACGTT	1740
GTTTGCTGTC	CATCATTGGT	GCTAGTTATG	GCCGAGCGGA	AGGATTATCG	AAGTGGTGGA	1800
CTTCGGGGCG	TTACCACCGG	AGATCAACTC	CGCGAGGATG	TACGCCGGCC	CGGGTTCGGC	1860
CTCGCTGGTG	GCCGCCGCGA	AGATGTGGGA	CAGCGTGGCG	AGTGACCTGT	TTTCGGCCGC	1920
GTCGGCGTTT	CAGTCGGTGG	TCTGGGGTCT	GACGACGGGA	TCGTGGATAG	GTTCGTCGGC	1980
GGGTCTGATG	GTGGCGGCGG	CCTCGCCGTA	TGTGGCGTGG	ATGAGCGTCA	CCGCGGGGCA	2040
GGCCGAGCTG	ACCGCCGCCC	AGGTCCGGGT	TGCTGCGGCG	GCCTACGAGA	CGGCGTATGG	2100
GCTGACGGTG	cccccccc	TGATCGCCGA	GAACCGTGCT	GAACTGATGA	TTCTGATAGC	2160
GACCAACCTC	TTGGGGCAAA	ACACCCCGGC	GATCGCGGTC	AACGAGGCCG	AATACGGGGA	2220
GATGTGGGCC	CAAGACGCCG	CCGCGATGTT	TGGCTACGCC	GCCACGGCGG	CGACGGCGAC	2280
CGAGGCGTTG	CTGCCGTTCG	AGGACGCCCC	ACTGATCACC	AACCCCGGCG	GGCTCCTTGA	2340
GCAGGCCGTC	GCGGTCGAGG	AGGCCATCGA	CACCGCCGCG	GCGAACCAGT	TGATGAACAA	2400
TGTGCCCCAA	GCGCTGCAAC	AACTGGCCCA	GCCCACGAAA	AGCATCTGGC	CGTTCGACCA	2460
ACTGAGTGAA	CTCTGGAAAG	CCATCTCGCC	GCATCTGTCG	CCGCTCAGCA	ACATCGTGTC	2520
GATGCTCAAC	AACCACGTGT	CGATGACCAA	CTCGGGTGTG	TCGATGGCCA	GCACCTTGCA	2580
CTCAATGTTG	AAGGGCTTTG	CTCCGGCGGC	GGCTCAGGCC	GTGGAAACCG	CGGCGCAAAA	2640
CGGGGTCCAG	GCGATGAGCT	CGCTGGGCAG	CCAGCTGGGT	TCGTCGCTGG	GTTCTTCGGG	2700
TCTGGGCGCT	GGGGTGGCCG	CCAACTTGGG	TCGGGCGGCC	TCGGTCGGTT	CGTTGTCGGT	276
GCCGCAGGCC	TGGGCCGCGG	CCAACCAGGC	GGTCACCCCG	GCGGCGCGG	CGCTGCCGCT	282
GACCAGCCTG	ACCAGCGCCG	CCCAAACCGC	CCCCGGACAC	ATGCTGGGCG	GGCTACCGCT	288
GGGGCAACTG	ACCAATAGCG	GCGGCGGGTT	CGGCGGGGTT	AGCAATGCGT	TGCGGATGCC	294
cccccccc	TACGTAATGC	CCCGTGTGCC	CGCCGCCGGG	TAACGCCGAT	CCGCACGCAA	300

TGCGGGCCCT CTATGCGGGC AGCGATC

3027

(2) INFORMATION FOR SEQ ID NO: 106:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 396 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:106:

Val Val Asp Phe Gly Ala Leu Pro Pro Glu Ile Asn Ser Ala Arg Met
1 5 10 15

Tyr Ala Gly Pro Gly Ser Ala Ser Leu Val Ala Ala Ala Lys Met Trp

Asp Ser Val Ala Ser Asp Leu Phe Ser Ala Ala Ser Ala Phe Gln Ser 35 40 45

Val Val Trp Gly Leu Thr Thr Gly Ser Trp Ile Gly Ser Ser Ala Gly 50 55 60

Leu Met Val Ala Ala Ala Ser Pro Tyr Val Ala Trp Met Ser Val Thr 65 70 75 80

Ala Gly Gln Ala Glu Leu Thr Ala Ala Gln Val Arg Val Ala Ala Ala 85 90 95

Ala Tyr Glu Thr Ala Tyr Gly Leu Thr Val Pro Pro Pro Val Ile Ala 100 105 110

Glu Asn Arg Ala Glu Leu Met Ile Leu Ile Ala Thr Asn Leu Leu Gly
115 120 125

Gln Asn Thr Pro Ala Ile Ala Val Asn Glu Ala Glu Tyr Gly Glu Met 130 135 140

Trp Ala Gln Asp Ala Ala Ala Met Phe Gly Tyr Ala Ala Thr Ala Ala 145 150 155 160

Thr Ala Thr Glu Ala Leu Leu Pro Phe Glu Asp Ala Pro Leu Ile Thr 165 170 175

Asn Pro Gly Gly Leu Leu Glu Gln Ala Val Ala Val Glu Glu Ala Ile 180 185 190

Asp Thr Ala Ala Ala Asn Gln Leu Met Asn Asn Val Pro Gln Ala Leu 195 200 205

Gln	Gln 210	Leu	Ala	Gln	Pro	Thr 215	Lys	Ser	Ile	Trp	Pro 220	Phe	Asp	Gln	Leu
Ser 225	Glu	Leu	Trp	Lys	Ala 230	Ile	Ser	Pro	His	Leu 235	Ser	Pro	Leu	Ser	Asn 240
Ile	Val	Ser	Met	Leu 245	Asn	Asn	His	Val	Ser 250	Met	Thr	Asn	Ser	Gly 255	Val
Ser	Met	Ala	Ser 260	Thr	Leu	His	Ser	Met 265	Leu	Lys	Gly	Phe	Ala 270	Pro	Ala
Ala	Ala	Gln 275	Ala	Val	Glu	Thr	Ala 280	Ala	Gln	Asn	Gly	Val 285	Gln	Ala	Met
Ser	Ser 290	Leu	Gly	Ser	Gln	Leu 295	Gly	Ser	Ser	Leu	Gly 300	Ser	Ser	Gly //	Leu
Gly 305	Ala	Gly	Val	Ala	Ala 310	Asn	Leu	Gly	Arg	Ala 315	Ala	Ser	Val	Gly	Ser 320
Leu	Ser	Val	Pro	Gln 325		Trp	Ala	Ala	Ala 330	Asn	Gln	Ala	Val	Thr 335	Pro
Ala	Ala	Arg	Ala 340	Leu	Pro	Leu	Thr	Ser 345	Leu	Thr	Ser	Ala	Ala 350	Gln	Thr
Ala	Pro	Gly 355	His	Met	Leu	Gly	Gly 360	Leu	Pro	Leu	Gly	Gln 365	Leu	Thr	Asn
Ser	Gly 370	Gly	Gly	Phe	Gly	Gly 375	Val	Ser	Asn	Ala	Leu 380	Arg	Met	Pro	Pro
Arg 385	Ala	Tyr	Val	Met	Pro 390	Arg	Val	Pro	Ala	Ala 395	Gly				

(2) INFORMATION FOR SEQ ID NO:107:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1616 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:107:

60	ATACCGCACG	ACCGGAGTAA	ACGCAATGCC	ATGCTGTGGC	GAGGG AGTGATCACC	CATCO
120	GGCAGACGCT	GCCGCGGGAT	GCTTGCGGCG	CGGCTCCAAT	TGGCC GGCGCGGTC	GCTGA
180	CTCTGGGAGA	CGCCTGAACT	GTTGACCGCG	AGGCCGTCGA	CCCCT CTGGACGCTC	ጥጥ ርርር

AGCCTGGACT	GGAGGTGGCA	GCGACAAGGC	GCTTGCGGCT	GCAACGCCGA	TGGTGGTCTG	240
CTACAAACC	GCGTCAACAC	AGGCCAAGAC	CCGTGCGATG	CAGGCGACGG	CGCAAGCCGC	300
GGCATACACC	CAGGCCATGG	CCACGACGCC	GTCGCTGCCG	GAGATCGCCG	CCAACCACAT	360
CACCCAGGCC	GTCCTTACGG	CCACCAACTT	CTTCGGTATC	AACACGATCC	CGATCGCGTT	420
GACCGAGATG	GATTATTTCA	TCCGTATGTG	GAACCAGGCA	GCCCTGGCAA	TGGAGGTCTA	480
CCAGGCCGAG	ACCGCGGTTA	ACACGCTTTT	CGAGAAGCTC	GAGCCGATGG	CGTCGATCCT	540
TGATCCCGGC	GCGAGCCAGA	GCACGACGAA	CCCGATCTTC	GGAATGCCCT	CCCCTGGCAG	600
CTCAACACCG	GTTGGCCAGT	TGCCGCCGGC	GGCTACCCAG	ACCCTCGGCC	AACTGGGTGA	660
GATGAGCGGC	CCGATGCAGC	AGCTGACCCA	GCCGCTGCAG	CAGGTGACGT	CGTTGTTCAG	720
CCAGGTGGGC	GGCACCGGCG	GCGGCAACCC	AGCCGACGAG	GAAGCCGCGC	AGATGGGCCT	780
GCTCGGCACC	AGTCCGCTGT	CGAACCATCC	GCTGGCTGGŢ	GGATCAGGCC	CCAGCGCGGG	840
CGCGGGCCTG	CTGCGCGCGG	AGTCGCTACC	TGGCGCAGGT	GGGTCGTTGA	CCCGCACGCC	900
GCTGATGTCT	CAGCTGATCG	AAAAGCCGGT	TGCCCCCTCG	GTGATGCCGG	CGGCTGCTGC	960
CGGATCGTCG	GCGACGGGTG	GCGCCGCTCC	GGTGGGTGCG	GGAGCGATGG	GCCAGGGTGC	1020
GCAATCCGGC	GGCTCCACCA	GGCCGGGTCT	GGTCGCGCCG	GCACCGCTCG	CGCAGGAGCG	1080
TGAAGAAGAC	GACGAGGACG	ACTGGGACGA	AGAGGACGAC	TGGTGAGCTC	CCGTAATGAC	1140
AACAGACTTC	CCGGCCACCC	GGGCCGGAAG	ACTTGCCAAC	ATTTTGGCG#	GGAAGGTAAA	1200
GAGAGAAAGT	AGTCCAGCAT	GGCAGAGATG	AAGACCGATG	CCGCTACCCT	CGCGCAGGAG	1260
					A GGTGGAGTCG	1320
ACGGCAGGTT	CGTTGCAGGG	CCAGTGGCGC	GGCGCGGCGG	GGACGGCCG	CCAGGCCGCG	1380
GTGGTGCGCT	TCCAAGAAGC	AGCCAATAAG	CAGAAGCAGG	AACTCGACG	A GATCTCGACG	1440
					A GCAGGCGCTG	
TCCTCGCAAA	TGGGCTTCTG	ACCCGCTAAT	ACGAAAAGA	A ACGGAGCAA	A AACATGACAG	1560
AGCAGCAGTG	GAATTTCGCG	GGTATCGAGG	CCGCGGCAAG	G CGCAATCCA	G GGAAAT	1616

(2) INFORMATION FOR SEQ ID NO:108:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 432 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single

TACGCCTCCG AA

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:108:

CTAGTGGATG GGACCATGGC CATTTCTGC AGTCTCACTG CCTTCTGTGT TGACATTTTG GCACGCCGGC GGAAACGAAG CACTGGGGTC GAAGAACGGC TGCGCTGCCA TATCGTCCGG

120

60

AGCTTCCATA CCTTCGTGCG GCCGGAAGAG CTTGTCGTAG TCGGCCGCCA TGACAACCTC

180

TCAGAGTGCG CTCAAACGTA TAAACACGAG AAAGGGCGAG ACCGACGGAA GGTCGAACTC

300

GCCCGATCCC GTGTTTCGCT ATTCTACGCG AACTCGGCGT TGCCCTATGC GAACATCCCA

360

GTGACGTTGC CTTCGGTCGA AGCCATTGCC TGACCGGCTT CGCTGATCGT CCGCGCCAGG

360

TTCTGCAGCG CGTTGTTCAG CTCGGTAGCC GTGGCGTCCC ATTTTTGCTG GACACCCTGG

420 432

(2) INFORMATION FOR SEQ ID NO:109:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 368 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:109:

Met Leu Trp His Ala Met Pro Pro Glu Xaa Asn Thr Ala Arg Leu Met
1 5 10 15

Ala Gly Ala Gly Pro Ala Pro Met Leu Ala Ala Ala Ala Gly Trp Gln 20 25 30

Thr Leu Ser Ala Ala Leu Asp Ala Gln Ala Val Glu Leu Thr Ala Arg 35 40 45

Leu Asn Ser Leu Gly Glu Ala Trp Thr Gly Gly Gly Ser Asp Lys Ala
50 55 60

Leu Ala Ala Ala Thr Pro Met Val Val Trp Leu Gln Thr Ala Ser Thr 65 70 75 80

Gln Ala Lys Thr Arg Ala Met Gln Ala Thr Ala Gln Ala Ala Tyr 85 90 95

Thr	Gln	Ala	Met 100	Ala	Thr	Thr	Pro	Ser 105	Leu	Pro	Glu	Ile	Ala 110	Ala	Asn
His	Ile	Thr 115	Gln	Ala	Val	Leu	Thr 120	Ala	Thr	Asn	Phe	Phe 125	Gly	Ile	Asn
Thr	Ile 130	Pro	Ile	Ala	Leu	Thr 135	Glu	Met	Asp	Tyr	Phe 140	Ile	Arg	Met	Trp
Asn 145	Gln	Ala	Ala	Leu	Ala 150	Met	Glu	Val	Tyr	Gln 155	Ala	Glu	Thr	Ala	Val 160
Asn	Thr	Leu	Phe	Glu 165	Lys	Leu	Glu	Pro	Met 170	Ala	Ser	Ile	Leu	Asp 175	Pro
Gly	Ala	Ser	Gln 180	Ser	Thr	Thr	Asn	Pro 185	Ile	Phe	Gly	Met	Pro 190	Ser	Pro
Gly	Ser	Ser 195	Thr	Pro	Val	Gly	Gln 200	Leu	Pro	Pro	Ala	Ala 205	Thr	Gĺn	Thr
Leu	Gly 210	Gln	Leu	Gly	Glu	Met 215	Ser	Gly	Pro	Met	Gln 220	Gln	Leu	Thr	Gln
Pro 225	Leu	Gln	Gln	Val	Thr 230	Ser	Leu	Phe	Ser	Gln 235	Val	Gly	Gly	Thr	Gly 240
Gly	Gly	Asn	Pro	Ala 245	Asp	Glu	Glu	Ala	Ala 250	Gln	Met	Gly	Leu	Leu 255	Gly
Thr	Ser	Pro	Leu 260	Ser	Asn	His	Pro	Leu 265	Ala	Gly	Gly	Ser	Gly 270	Pro	Ser
Ala	Gly	Ala 275	Gly	Leu	Leu	Arg	Ala 280	Glu	Ser	Leu	Pro	Gly 285	Ala	Gly	Gly
Ser	Leu 290	Thr	Arg	Thr	Pro	Leu 295	Met	Ser	Gln	Leu	11e 300	Glu	Lys	Pro	Val
Ala 305	Pro	Ser	Val	Met	Pro 310	Ala	Ala	Ala	Ala	Gly 315	Ser	Ser	Ala	Thr	Gly 320
Gly	Ala	Ala	Pro	Val 325	Gly	Ala	Gly	Ala	Met 330	Gly	Gln	Gly	Ala	Gln 335	Ser
Gly	Gly	Ser	Thr 340	Arg	Pro	Gly	Leu	Val 345		Pro	Ala	Pro	Leu 350	Ala	Gln
Glu	Arg	Glu 355	Glu	Asp	Asp	Glu	Asp 360		Trp	Asp	Glu	Glu 365	Asp	Asp	Trp

⁽²⁾ INFORMATION FOR SEQ ID NO:110:

133	SEQUENCE	CHARACTERISTICS:

(A) LENGTH: 100 amino acids

(B) TYPE: amino acid

(C) STRANDEDNESS:

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:110:

Met Ala Glu Met Lys Thr Asp Ala Ala Thr Leu Ala Gln Glu Ala Gly
1 5 10 15

Asn Phe Glu Arg Ile Ser Gly Asp Leu Lys Thr Gln Ile Asp Gln Val 20 25 30

Glu Ser Thr Ala Gly Ser Leu Gln Gly Gln Trp Arg Gly Ala Ala Gly 35 40 45

Thr Ala Ala Gln Ala Ala Val Val Arg Phe Gln Glu Ala Ala Asn Lys 50 55 60

Gln Lys Gln Glu Leu Asp Glu Ile Ser Thr Asn Ile Arg Gln Ala Gly 65 70 75 80

Val Gln Tyr Ser Arg Ala Asp Glu Glu Gln Gln Gln Ala Leu Ser Ser 85 90 95

Gln Met Gly Phe 100

(2) INFORMATION FOR SEQ ID NO:111:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 396 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:111:

60	GTTCGTTGCA	TCGACGGCAG	CCAGGTGGAG	CCCAGATCGA	GACCTGAAAA	GATCTCCGGC
120	GCTTCCAAGA	GCGGTGGTGC	CGCCCAGGCC	CGGGGACGGC	CGCGGCGCGG	GGGCCAGTGG
180	GTCAGGCCGG	ACGAATATTC	CGAGATCTCG	AGGAACTCGA	AAGCAGAAGC	AGCAGCCAAT
240	AAATGGGCTT	CTGTCCTCGC	GCAGCAGGCG	ACGAGGAGCA	TCGAGGGCCG	ССТССАВТАС

			1.0			
CTGACCCGCT	AATACGAAAA	GAAACGGAGC	AAAAACATGA	CAGAGCAGCA	GTGGAATTTC	300
GCGGGTATCG	AGGCCGCGC	AAGCGCAATC	CAGGGAAATG	TCACGTCCAT	TCATTCCCTC	360
CTTGACGAGG	GGAAGCAGTC	CCTGACCAAG	CTCGCA			396
(2) INFORM	ATION FOR SE	CQ ID NO:112	2:			
((A) LENGTH: B) TYPE: am	NESS: singl	cids			

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:112:

Ile Ser Gly Asp Leu Lys Thr Gln Ile Asp Gln Val Glu Ser Thr Ala 1 5 10 15

Gly Ser Leu Gln Gly Gln Trp Arg Gly Ala Ala Gly Thr Ala Ala Gln 20 25 30

Ala Ala Val Val Arg Phe Gln Glu Ala Ala Asn Lys Gln Lys Gln Glu 35 40 45

Leu Asp Glu Ile Ser Thr Asn Ile Arg Gln Ala Gly Val Gln Tyr Ser 50 55 60

Arg Ala Asp Glu Glu Gln Gln Gln Ala Leu Ser Ser Gln Met Gly Phe 65 70 75 80

(2) INFORMATION FOR SEQ ID NO:113:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 387 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:113:

GTGGATCCCG	ATCCCGTGTT	TCGCTATTCT	ACGCGAACTC	GGCGTTGCCC	TATGCGAACA	60
TCCCAGTGAC	GTTGCCTTCG	GTCGAAGCCA	TTGCCTGACC	GGCTTCGCTG	ATCGTCCGCG	120
CCAGGTTCTG	CAGCGCGTTG	TTCAGCTCGG	TAGCCGTGGC	GTCCCATTTT	TGCTGGACAC	180

CCTGGTACGC	CTCCGAACCG	CTACCGCCCC	AGGCCGCTGC	GAGCTTGGTC	AGGGACTGCT	240
TCCCCTCGTC	AAGGAGGGAA	TGAATGGACG	TGACATTTCC	CTGGATTGCG	CTTGCCGCGG	300
CCTCGATACC	CGCGAAATTC	CACTGCTGCT	CTGTCATGTT	TTTGCTCCGT	TTCTTTTCGT	360
ATTAGCGGGT	CAGAAGCCCA	TTTGCGA				387

(2) INFORMATION FOR SEQ ID NO:114:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 272 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:114:

CGGCACGAGG ATCTCGGTTG GCCCAACGGC GCTGGCGAGG GCTCCGTTCC GGGGGCGAGC 60

TGCGCGCCGG ATGCTTCCTC TGCCCGCAGC CGCGCCTGGA TGGATGGACC AGTTGCTACC 120

TTCCCGACGT TTCGTTCGGT GTCTGTGCGA TAGCGGTGAC CCCGGCGCGC ACGTCGGAGG 180

TGTTGGGGGG CAGGCCGGGT CGGTGGTTCG GCCGGGACG CAGACGGTCT GGACGGAACG 240

GGCGGGGGTT CGCCGATTGG CATCTTTGCC CA 272

- (2) INFORMATION FOR SEQ ID NO:115:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 20 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:115:

Asp Pro Val Asp Ala Val Ile Asn Thr Thr Cys Asn Tyr Gly Gln Val 1 5 10 15

Val Ala Ala Leu 20

(2) INFORMATION FOR SEQ ID NO:116:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 15 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:116:

Ala Val Glu Ser Gly Met Leu Ala Leu Gly Thr Pro Ala Pro Ser

1 5 10 15

- (2) INFORMATION FOR SEQ ID NO:117:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:117:

Ala Ala Met Lys Pro Arg Thr Gly Asp Gly Pro Leu Glu Ala Ala Lys

1 10 15

Glu Gly Arg

- (2) INFORMATION FOR SEQ ID NO:118:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 15 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:118:

Tyr Tyr Trp Cys Pro Gly Gln Pro Phe Asp Pro Ala Trp Gly Pro
1 5 10 15

- (2) INFORMATION FOR SEQ ID NO:119:
 - (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 14 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:119:

Asp Ile Gly Ser Glu Ser Thr Glu Asp Gln Gln Xaa Ala Val

- (2) INFORMATION FOR SEQ ID NO:120:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 13 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:120:

Ala Glu Glu Ser Ile Ser Thr Xaa Glu Xaa Ile Val Pro 1 5 10

- (2) INFORMATION FOR SEQ ID NO:121:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 17 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:121:

Asp Pro Glu Pro Ala Pro Pro Val Pro Thr Thr Ala Ala Ser Pro Pro 1 5 10 15

Ser

- (2) INFORMATION FOR SEQ ID NO:122:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 15 amino acids

- (B) TYPE: amino acid
- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:122:

Ala Pro Lys Thr Tyr Xaa Glu Glu Leu Lys Gly Thr Asp Thr Gly
1 5 10 15

- (2) INFORMATION FOR SEQ ID NO:123:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 30 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:123:

Asp Pro Ala Ser Ala Pro Asp Val Pro Thr Ala Ala Gln Leu Thr Ser 1 5 10 15

Leu Leu Asn Ser Leu Ala Asp Pro Asn Val Ser Phe Ala Asn 20 25 30

- (2) INFORMATION FOR SEQ ID NO:124:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 22 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:124:

Asp Pro Pro Asp Pro His Gln Xaa Asp Met Thr Lys Gly Tyr Tyr Pro 1 5 10 15

Gly Gly Arg Arg Xaa Phe 20

(2) INFORMATION FOR SEQ ID NO:125:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 7 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:125:

Asp Pro Gly Tyr Thr Pro Gly 5

- (2) INFORMATION FOR SEQ ID NO:126:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 10 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ix) FEATURE: (D) OTHER INFORMATION: /note= "The Second Residue Can Be Either a Pro or Thr"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: :

Xaa Xaa Gly Phe Thr Gly Pro Gln Tyr

- (2) INFORMATION FOR SEQ ID NO:127:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 9 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ix) FEATURE:

(D) OTHER INFORMATION: /note= "The Third Residue Can Be Either a Gln or Leu"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:127:

Xaa Pro Xaa Val Thr Ala Tyr Ala Gly 5

- (2) INFORMATION FOR SEQ ID NO:128:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 9 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:128:

Xaa Xaa Xaa Glu Lys Pro Phe Leu Arg 1 5

- (2) INFORMATION FOR SEQ ID NO:129:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 15 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:129:

- (2) INFORMATION FOR SEQ ID NO:130:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 15 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:130:

Ala Gly Asp Thr Xaa Ile Tyr Ile Val Gly Asn Leu Thr Ala Asp 1 5 10 15

- (2) INFORMATION FOR SEQ ID NO:131:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 15 amino acids

(B)	TYPE:	amino	acid
\	OMBANI	20011000	٠.

(C) STRANDEDNESS:

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:131:

Ala Pro Glu Ser Gly Ala Gly Leu Gly Gly Thr Val Gln Ala Gly
1 5 10 15

- (2) INFORMATION FOR SEQ ID NO:132:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 21 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:132:

Xaa Tyr Ile Ala Tyr Xaa Thr Thr Ala Gly Ile Val Pro Gly Lys Ile 1 5 10 15

Asn Val His Leu Val 20

- (2) INFORMATION FOR SEQ ID NO:133:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 882 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: DNA (genomic)
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:133:

GCAACGCTGT CGTGGCCTTT GCGGTGATCG GTTTCGCCTC GCTGGCGGT GCGGTGGCGG 60

TCACCATCCG ACCGACCGCG GCCTCAAAAC CGGTAGAGGG ACACCAAAAC GCCCAGCCAG 120

GGAAGTTCAT GCCGTTGTTG CCGACGCAAC AGCAGGCGCC GGTCCCGCCG CCTCCGCCCG 180

ATGATCCCAC CGCTGGATTC CAGGGCGGCA CCATTCCGGC TGTACAGAAC GTGGTGCCGC 240

GGCCGGGTAC	CTCACCCGGG	GTGGGTGGGA	CGCCGGCTTC	GCCTGCGCCG	GAAGCGCCGG	300
CCGTGCCCGG	TGTTGTGCCT	GCCCCGGTGC	CAATCCCGGT	CCCGATCATC	ATTCCCCCGT	360
TCCCGGGTTG	GCAGCCTGGA	ATGCCGACCA	TCCCCACCGC	ACCGCCGACG	ACGCCGGTGA	420
CCACGTCGGC	GACGACGCCG	CCGACCACGC	CGCCGACCAC	GCCGGTGACC	ACGCCGCCAA	480
CGACGCCGCC	GACCACGCCG	GTGACCACGC	CGCCAACGAC	GCCGCCGACC	ACGCCGGTGA	540
CCACGCCACC	AACGACCGTC	GCCCCGACGA	CCGTCGCCCC	GACGACGGTC	GCTCCGACCA	600
CCGTCGCCCC	GACCACGGTC	GCTCCAGCCA	CCGCCACGCC	GACGACCGTC	GCTCCGCAGC	660
CGACGCAGCA	GCCCACGCAA	CAACCAACCC	AACAGATGCC	AACCCAGCAG	CAGACCGTGG	720
CCCCGCAGAC	GGTGGCGCCG	GCTCCGCAGC	CGCCGTCCGG	TGGCCGCAAC	GGCAGCGGCG	780
GGGCGACTT	ATTCGGCGGG	TTCTGATCAC	GGTCGCGGCT	TCACTACGGT	CGGAGGACAT	840
GGCCGGTGAT	GCGGTGACGG	TGGTGCTGCC	CTGTCTCAAC	GA		882

(2) INFORMATION FOR SEQ ID NO:134:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 815 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:134:

CCATCAACCA	ACCGCTCGCG	CCGCCCGCGC	CGCCGGATCC	GCCGTCGCCG	CCACGCCCGC	60
CGGTGCCTCC	GGTGCCCCCG	TTGCCGCCGT	CGCCGCCGTC	GCCGCCGACC	GGCTGGGTGC	120
CTAGGGCGCT	GTTACCGCCC	TGGTTGGCGG	GGACGCCGCC	GGCACCACCG	GTACCGCCGA	180
TGGCGCCGTT	GCCGCCGGCG	GCACCGTTGC	CACCGTTGCC	ACCGTTGCCA	CCGTTGCCGA	240
CCAGCCACCC	GCCGCGACCA	CCGGCACCGC	CGGCGCCGCC	CGCACCGCCG	GCGTGCCCGT	300
TCGTGCCCGT	ACCGCCGGCA	CCGCCGTTGC	CGCCGTCACC	GCCGACGGAA	CTACCGGCGG	360
ACGCGGCCTG	CCCGCCGGCG	CCGCCCGCAC	CGCCATTGGC	ACCGCCGTCA	CCGCCGGCTG	420
GGAGTGCCGC	GATTAGGGCA	CTGACCGGCG	CAACCAGCGC	AAGTACTCTC	GGTCACCGAG	480
CACTTCCAGA	CGACACCACA	GCACGGGGTT	GTCGGCGGAC	TGGGTGAAAT	GGCAGCCGAT	540

AGCGGCTAGC	TGTCGGCTGC	GGTCAACCTC	GATCATGATG	TCGAGGTGAC	CGTGACCGCG	600
CCCCCGAAG	GAGGCGCTGA	ACTCGGCGTT	GAGCCGATCG	GCGATCGGTT	GGGGCAGTGC	660
CCAGGCCAAT	ACGGGGATAC	CGGGTGTCNA	AGCCGCCGCG	AGCGCAGCTT	CGGTTGCGCG	720
ACNGTGGTCG	GGGTGGCCTG	TTACGCCGTT	GTCNTCGAAC	ACGAGTAGCA	GGTCTGCTCC	780
GGCGAGGGCA	TCCACCACGC	GTTGCGTCAG	CTCGT			815

(2) INFORMATION FOR SEQ ID NO:135:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1152 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:135:

ACCAGCCGCC	GGCTGAGGTC	TCAGATCAGA	GAGTCTCCGG	ACTCACCGGG	GCGGTTCAGC	60
CTTCTCCCAG	AACAACTGCT	GAAGATCCTC	GCCCGCGAAA	CAGGCGCTGA	TTTGACGCTC	120
TATGACCGGT	TGAACGACGA	GATCATCCGG	CAGATTGATA	TGGCACCGCT	GGGCTAACAG	180
GTGCGCAAGA	TGGTGCAGCT	GTATGTCTCG	GACTCCGTGT	CGCGGATCAG	CTTTGCCGAC	240
GGCCGGGTGA	TCGTGTGGAG	CGAGGAGCTC	GGCGAGAGCC	AGTATCCGAT	CGAGACGCTG	300
GACGGCATCA	CGCTGTTTGG	GCGGCCGACG	ATGACAACGC	CCTTCATCGT	TGAGATGCTC	360
AAGCGTGAGC	GCGACATCCA	GCTCTTCACG	ACCGACGGCC	ACTACCAGGG	CCGGATCTCA	420
ACACCCGACG	TGTCATACGC	GCCGCGGCTC	CGTCAGCAAG	TTCACCGCAC	CGACGATCCT	480
GCGTTCTGCC	TGTCGTTAAG	CAAGCGGATC	GTGTCGAGGA	AGATCCTGAA	TCAGCAGGCC	540
TTGATTCGGG	CACACACGTC	GGGGCAAGAC	GTTGCTGAGA	GCATCCGCAC	GATGAAGCAC	600
TCGCTGGCCT	GGGTCGATCG	ATCGGGCTCC	CTGGCGGAGT	TGAACGGGTT	CGAGGGAAAT	660
GCCGCAAAGG	CATACTTCAC	CGCGCTGGGG	CATCTCGTCC	CGCAGGAGTT	CGCATTCCAG	720
GGCCGCTCGA	CTCGGCCGCC	GTTGGACGCC	TTCAACTCGA	TGGTCAGCCT	CGGCTATTCG	780
CTGCTGTACA	AGAACATCAT	AGGGGCGATC	GAGCGTCACA	GCCTGAACGC	GTATATCGGT	840
TTCCTACACC	AGGATTCACG	AGGGCACGCA	ACGTCTCGTG	CCGAATTCGG	CACGAGCTCC	900

GCTGAAACCG	CTGGCCGGCT	GCTCAGTGCC	CGTACGTAAT	CCGCTGCGCC	CAGGCCGGCC	960
CGCCGGCCGA	ATACCAGCAG	ATCGGACAGC	GAATTGCCGC	CCAGCCGGTT	GGAGCCGTGC	1020
ATACCGCCGG	CACACTCACC	GGCAGCGAAC	AGGCCTGGCA	CCGTGGCGGC	GCCGGTGTCC	1080
GCGTCTACTT	CGACACCGCC	CATCACGTAG	TGACACGTCG	GCCCGACTTC	CATTGCCTGC	1140
GTTCGGCACG	AG					1152

(2) INFORMATION FOR SEQ ID NO:136:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 655 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:136:

CTCGTGCCG	TTCGGCAGGG	TGTACTTGCC	GGTGGTGTAN	GCCGCATGAG	TGCCGACGAC	6 Ó
CAGCAATGC	GCAACAGCAC	GGATCCCGGT	CAACGACGCC	ACCCGGTCCA	CGTGGGCGAT	120
CCGCTCGAGT	CCGCCCTGGG	CGGCTCTTTC	CTTGGGCAGG	GTCATCCGAC	GTGTTTCCGC	180
CGTGGTTTGC	CGCCATTATG	CCGGCGCGCC	GCGTCGGGCG	GCCGGTATGG	CCGAANGTCG	240
ATCAGCACAC	CCGAGATACG	GGTCTGTGCA	AGCTTTTTGA	GCGTCGCGCG	GGGCAGCTTC	300
GCCGGCAATT	CTACTAGCGA	GAAGTCTGGC	CCGATACGGA	TCTGACCGAA	GTCGCTGCGG	360
TGCAGCCCAC	CCTCATTGGC	GATGGCGCCG	ACGATGGCGC	CTGGACCGAT	CTTGTGCCGC	420
TTGCCGACGG	CGACGCGGTA	GGTGGTCAAG	TCCGGTCTAC	GCTTGGGCCT	TTGCGGACGG	480
TCCCGACGCT	GGTCGCGGTT	GCGCCGCGAA	AGCGGCGGGT	CGGGTGCCAT	CAGGAATGCC	540
TCACCGCCGC	GGCACTGCAC	GGCCAGTGCC	GCGGCGATGT	CAGCCATCGG	GACATCATGC	600
TCGCGTTCAT	ACTCCTCGAC	CAGTCGGCGG	AACAGCTCGA	TTCCCGGACC	GCCCA	655

(2) INFORMATION FOR SEQ ID NO:137:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 267 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(xi)	SEQ	JENCI	E DES	SCRII	OITS	1: SI	EQ II	NO:	137:	:					
Asn 1	Ala	Val	Val	Ala 5	Phe	Ala	Val	Ile	Gly 10	Phe	Ala	Ser	Leu	Ala 15	Val
Ala	Val	Ala	Val 20	Thr	Ile	Arg	Pro	Thr 25	Ala	Ala	Ser	Lys	Pro 30	Val	Glu
Gly	His	Gln 35	Asn	Ala	Gln	Pro	Gly 40	Lys	Phe	Met	Pro	Leu 45	Leu	Pro	Thr
Gln	Gln 50	Gln	Ala	Pro	Val	Pro 55	Pro	Pro	Pro	Pro	Asp 60	Asp	Pro	Thr	Ala
Gly 65	Phe	Gln	Gly	Gly	Thr 70	Ile	Pro	Ala	Val	Gln 75	Asn	Val	Val	Pro	Arg 80
Pro	Gly	Thr	Ser	Pro 85	Gly	Val	Gly	Gly	Thr 90	Pro	Ala	Ser	Pro	Ala 95	Pro
Glu	Ala	Pro	Ala 100	Val	Pro	Gly	Val	Val 105	Pro	Ala	Pro	Val	Pro 110	Ile	Pro
Val	Pro	Ile 115	Ile	Ile	Pro	Pro	Phe 120	Pro	Gly	Trp	Gln	Pro 125	Gly	Met	Pro
Thr	Ile 130	Pro	Thr	Ala	Pro	Pro 135	Thr	Thr	Pro	Val	Thr 140	Thr	Ser	Ala	Thr
Thr 145	Pro	Pro	Thr	Thr	Pro 150	Pro	Thr	Thr	Pro	Val 155	Thr	Thr	Pro	Pro	Thr 160
Thr	Pro	Pro	Thr	Thr 165	Pro	Val	Thr	Thr	Pro 170	Pro	Thr	Thr	Pro	Pro 175	Thr
Thr	Pro	Val	Thr 180	Thr	Pro	Pro	Thr	Thr 185	Val	Ala	Pro	Thr	Thr 190	Val	Ala
Pro	Thr	Thr 195	Val	Ala	Pro	Thr			Ala		Thr	Thr 205		Ala	Pro
Ala	Thr 210	Ala	Thr	Pro	Thr	Thr 215	Val	Ala	Pro	Gln	Pro 220	Thr	Gln	Gln	Pro
Thr 225		Gln	Pro	Thr	Gln 230	Gln	Met	Pro	Thr	Gln 235	Gln	Gln	Thr	Val	Ala 240
Pro	Gln	Thr	Val	Ala 245	Pro	Ala	Pro	Gln	Pro 250	Pro	Ser	Gly	Gly	Arg 255	Asn

Gly Ser Gly Gly Gly Asp Leu Phe Gly Gly Phe 260 265

- (2) INFORMATION FOR SEQ ID NO:138:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 174 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:138:
 - Ile Asn Gln Pro Leu Ala Pro Pro Ala Pro Pro Asp Pro Pro Ser Pro 1 5 10 15
 - Pro Arg Pro Pro Val Pro Pro Val Pro Pro Leu Pro Pro Ser Pro Pro 20 25 30
 - Ser Pro Pro Thr Gly Trp Val Pro Arg Ala Leu Leu Pro Pro Trp Leu 35 40 45
 - Ala Gly Thr Pro Pro Ala Pro Pro Val Pro Pro Met Ala Pro Leu Pro 50 55 60
 - Pro Ala Ala Pro Leu Pro Thr 65
 - Ser His Pro Pro Arg Pro Pro Ala Pro Pro Ala Pro Pro Ala Pro Pro 85 90 95
 - Ala Cys Pro Phe Val Pro Val Pro Pro Ala Pro Pro Leu Pro Pro Ser 100 105 110
 - Pro Pro Thr Glu Leu Pro Ala Asp Ala Ala Cys Pro Pro Ala Pro Pro 115 120 125
 - Ala Pro Pro Leu Ala Pro Pro Ser Pro Pro Ala Gly Ser Ala Ala Ile 130 135 140
 - Arg Ala Leu Thr Gly Ala Thr Ser Ala Ser Thr Leu Gly His Arg Ala 145 150 155 160
 - Leu Pro Asp Asp Thr Thr Ala Arg Gly Cys Arg Arg Thr Gly 165 170
- (2) INFORMATION FOR SEQ ID NO:139:
 - (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 35 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:139:

Gln Pro Pro Ala Glu Val Ser Asp Gln Arg Val Ser Gly Leu Thr Gly
1 5 10 15

Ala Val Gln Pro Ser Pro Arg Thr Thr Ala Glu Asp Pro Arg Pro Arg 20 25 30

Asn Arg Arg 35

- (2) INFORMATION FOR SEQ ID NO:140:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 104 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:140:

Arg Ala Asp Ser Ala Gly Cys Thr Cys Arg Trp Cys Xaa Pro His Glu 1 5 10 15

Cys Arg Arg Pro Ala Met Arg Gln Gln His Gly Ser Arg Ser Thr Thr 20 25 30

Pro Pro Gly Pro Arg Gly Arg Ser Ala Arg Val Arg Pro Gly Arg Leu 35 40 45

Phe Pro Trp Ala Gly Ser Ser Asp Val Phe Pro Pro Trp Phe Ala Ala 50 55 60

Ile Met Pro Ala Arg Arg Val Gly Arg Pro Val Trp Pro Xaa Val Asp
65 70 75 80

Gln His Thr Arg Asp Thr Gly Leu Cys Lys Leu Phe Glu Arg Arg Ala 85 90 95

Gly Gln Leu Arg Arg Gln Phe Tyr

(2) INFORMATION FOR SEQ ID NO:141:
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 53 base pairs (B) TYPE: nucleic acid
(C) STRANDEDNESS: single(D) TOPOLOGY: linear
(ii) MOLECULE TYPE: other nucleic acid

- (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Mycobacterium tuberculosis

(A) DESCRIPTION: /desc = "PCR primer"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:141:

GGATCCATAT GGGCCATCAT CATCATCATC ACGTGATCGA CATCATCGGG ACC

53

- (2) INFORMATION FOR SEQ ID NO:142:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 42 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Mycobacterium tuberculosis
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:142: CCTGAATTCA GGCCTCGGTT GCGCCGGCCT CATCTTGAAC GA

42

- (2) INFORMATION FOR SEQ ID NO:143:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 31 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Mycobacterium tuberculosis

	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:143:	
GGA:	TCCTGCA GGCTCGAAAC CACCGAGCGG T	31
(2)	INFORMATION FOR SEQ ID NO:144:	
	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 31 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
٠	(ii) MOLECULE TYPE: other nucleic acid (A) DESCRIPTION: /desc = "PCR primer" /	
	(vi) ORIGINAL SOURCE: (A) ORGANISM: Mycobacterium tuberculosis	÷
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:144:	
CTC	TGAATTC AGCGCTGGAA ATCGTCGCGA T	31
(2)	INFORMATION FOR SEQ ID NO:145:	
	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 33 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
	<pre>(ii) MOLECULE TYPE: other nucleic acid (A) DESCRIPTION: /desc = "PCR primer"</pre>	
	(vi) ORIGINAL SOURCE: (A) ORGANISM: Mycobacterium tuberculosis	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:145:	
GGA	TCCAGCG CTGAGATGAA GACCGATGCC GCT	33
(2)	INFORMATION FOR SEQ ID NO:146:	
	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 33 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	

<pre>(ii) MOLECULE TYPE: other nucleic acid (A) DESCRIPTION: /desc = "PCR primer"</pre>	
<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Mycobacterium tuberculosis</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:146:	
GAGAGAATTC TCAGAAGCCC ATTTGCGAGG ACA	33
(2) INFORMATION FOR SEQ ID NO:147:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1993 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(ii) MOLECULE TYPE: DNA (genomic)	
<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Mycobacterium tuberculosis</pre>	
(ix) FEATURE: (A) NAME/KEY: CDS (B) LOCATION: 1521273	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:147:	
TGTTCTTCGA CGGCAGGCTG GTGGAGGAAG GGCCCACCGA ACAGCTGTTC TCCTCGCCGA	60
AGCATGCGGA AACCGCCCGA TACGTCGCCG GACTGTCGGG GGACGTCAAG GACGCCAAGC	120
GCGGAAATTG AAGAGCACAG AAAGGTATGG C GTG AAA ATT CGT TTG CAT ACG Val Lys Ile Arg Leu His Thr 1 5	172
CTG TTG GCC GTG TTG ACC GCT GCG CCG CTG CTG CTA GCA GCG GCG GGC Leu Leu Ala Val Leu Thr Ala Ala Pro Leu Leu Leu Ala Ala Ala Gly 10 15 20	220
TGT GGC TCG AAA CCA CCG AGC GGT TCG CCT GAA ACG GGC GCC GGC GCC Cys Gly Ser Lys Pro Pro Ser Gly Ser Pro Glu Thr Gly Ala Gly Ala 25	268
GGT ACT GTC GCG ACT ACC CCC GCG TCG TCG CCG GTG ACG TTG GCG GAG Gly Thr Val Ala Thr Thr Pro Ala Ser Ser Pro Val Thr Leu Ala Glu 40 45 50 55	316
ACC GGT AGC ACG CTG CTC TAC CCG CTG TTC AAC CTG TGG GGT CCG GCC Thr Gly Ser Thr Leu Leu Tyr Pro Leu Phe Asn Leu Trp Gly Pro Ala 60 65 70	364

TTT	CAC His	GAG Glu	AGG Arg 75	TAT Tyr	CCG Pro	AAC Asn	GTC Val	ACG Thr 80	ATC Ile	ACC Thr	GCT Ala	CAG Gln	GGC Gly 85	ACC Thr	GGT Gly		412
TCI	GGT Gly	GCC Ala 90	GGG Gly	ATC Ile	GCG Ala	CAG Gln	GCC Ala 95	GCC Ala	GCC Ala	GGG Gly	ACG Thr	GTC Val 100	AAC Asn	ATT Ile	GGG Gly		460
GCC Ala	TCC Ser 105	GAC Asp	GCC Ala	TAT Tyr	CTG Leu	TCG Ser 110	GAA Glu	GGT Gly	GAT Asp	ATG Met	GCC Ala 115	GCG Ala	CAC His	AAG Lys	GGG Gly		508
CTC Lev	ATG Met	AAC Asn	ATC Ile	GCG Ala	CTA Leu 125	GCC Ala	ATC Ile	TCC Ser	GCT Ala	CAG Gln 130	CAG Gln	GTC Val	AAC Asn	TAC Tyr	AAC Asn 135	,	556
CTC	CCC Pro	GGA Gly	GTG Val	AGC Ser 140	GAG Glu	CAC His	CTC Leu	AAG Lys	CTG Leu 145	AAC Asn	GGA Gly	AAA Lys	GTC Val	CTG Leu 150	GCG Ala		604
GCC Ala	ATG Met	TAC Tyr	CAG Gln 155	GGC Gly	ACC Thr	ATC Ile	AAA Lys	ACC Thr 160	TGG Trp	GAC Asp	GAC Asp	CCG Pro	CAG Gln 165	ATC Ile	GCT Ala		652
GC0 Ala	CTC Leu	AAC Asn 170	CCC Pro	GGC Gly	GTG Val	AAC Asn	CTG Leu 175	CCC Pro	GGC Gly	ACC Thr	GCG Ala	GTA Val 180	GTT Val	CCG Pro	CTG Leu		700
CAC His	CGC Arg 185	TCC Ser	GAC Asp	GGG Gly	TCC Ser	GGT Gly 190	GAC Asp	ACC Thr	TTC Phe	TTG Leu	TTC Phe 195	ACC Thr	CAG Gln	TAC Tyr	CTG Leu		748
TCC Ser 200	AAG Lys	CAA Gln	GAT Asp	CCC Pro	GAG Glu 205	GGC Gly	TGG Trp	GGC Gly	AAG Lys	TCG Ser 210	CCC Pro	GGC Gly	TTC Phe	GGC Gly	ACC Thr 215		796
ACC Thi	GTC Val	GAC Asp	TTC Phe	CCG Pro 220	GCG Ala	GTG Val	CCG Pro	GGT Gly	GCG Ala 225	CTG Leu	GGT Gly	GAG Glu	AAC Asn	GGC Gly 230	AAC Asn	:	844
GG(Gl _y	GGC Gly	ATG Met	GTG Val 235	ACC Thr	GGT Gly	TGC Cys	GCC Ala	GAG Glu 240	ACA Thr	CCG Pro	GGC Gly	TGC Cys	GTG Val 245	GCC Ala	TAT Tyr		892
ATC Ile	GGC Gly	ATC Ile 250	AGC Ser	TTC Phe	CTC Leu	GAC Asp	CAG Gln 255	GCC Ala	AGT Ser	CAA Gln	CGG Arg	GGA Gly 260	Leu	G1 y	GAG Glu		940
Alá	Gln 265	Leu	Gly	Asn	Ser	Ser 270	Gly	Asn	Phe	Leu	Leu 275	Pro	Asp	Ala			988
AG(Se)	ATT Ile	CAG Gln	GCC Ala	GCG Ala	GCG Ala	GCT Ala	GGC Gly	TTC Phe	GCA Ala	TCG Ser	AAA Lys	ACC Thr	CCG Pro	GCG Ala	AAC Asn		1036

280	285	29	0	295
CAG GCG ATT TCG AT Gln Ala Ile Ser Me 30	t Ile Asp Gly	CCC GCC CC Pro Ala Pr 305	G GAC GGC TAC o Asp Gly Tyr	CCG ATC 1084 Pro Ile 310
ATC AAC TAC GAG TA Ile Asn Tyr Glu Ty 315	C GCC ATC GTC Ala Ile Val	AAC AAC CG Asn Asn Arc 320	G CAA AAG GAC g Gln Lys Asp 325	GCC GCC 1132 Ala Ala
ACC GCG CAG ACC TTO Thr Ala Gln Thr Lev 330	G CAG GCA TTT Gln Ala Phe 335	CTG CAC TGG Leu His Trp	G GCG ATC ACC P Ala Ile Thr 340	GAC GGC 1180 Asp Gly
AAC AAG GCC TCG TTC Asn Lys Ala Ser Phe 345	CTC GAC CAG Leu Asp Gln 350	GTT CAT TTO Val His Phe	C CAG CCG CTG e Gln Pro Leu 355	CCG CCC 1228 Pro Pro
GCG GTG GTG AAG TTG Ala Val Val Lys Leu 360	TCT GAC GCG Ser Asp Ala 365	TTG ATC GCG Leu Ile Ala 370	Thr Ile Ser	AGC / 1273 Ser /
TAGCCTCGTT GACCACCA	CG CGACAGCAAC	CTCCGTCGGG	CCATCGGGCT G	CTTTGCGGA 1333
GCATGCTGGC CCGTGCCG	GT GAAGTCGGCC	GCGCTGGCCC	GGCCATCCGG TO	GGTTGGGTG 1393
GGATAGGTGC GGTGATCC	CG CTGCTTGCGC	TGGTCTTGGT	GCTGGTGGTG C	rggtcatcg 1453
AGGCGATGGG TGCGATCA	GG CTCAACGGGT	TGCATTTCTT	CACCGCCACC GA	AATGGAATC 1513
CAGGCAACAC CTACGGCG			•	· · · · ·
CTACGGGGCG TTGCCGCTC				
CGCGGTGCCG GTCTCTGTA				
GGCCGAGGCT GTGGGAATA				
TTTGTGGGGG GCAATGACG				
TCACAACGCT CCCGATGTG				
GGGCATGTTG GTGTCCGGT				
CACTCATGAC CTGTTCCGG	C AGGTGCCGGT	GTTGCCCCGG	GAGGGCGCGA TC	GGGAATTC 1993

(2) INFORMATION FOR SEQ ID NO:148:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 374 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:148:
- Val Lys Ile Arg Leu His Thr Leu Leu Ala Val Leu Thr Ala Ala Pro
- Leu Leu Leu Ala Ala Ala Gly Cys Gly Ser Lys Pro Pro Ser Gly Ser 20 25 30
- Pro Glu Thr Gly Ala Gly Ala Gly Thr Val Ala Thr Thr Pro Ala Ser 35 40 45
- Ser Pro Val Thr Leu Ala Glu Thr Gly Ser Thr Leu Leu Tyr Pro Leu 50 55 60
- Phe Asn Leu Trp Gly Pro Ala Phe His Glu Arg Tyr Pro Asn Val Thr 65 70 75 80
- Ile Thr Ala Gln Gly Thr Gly Ser Gly Ala Gly Ile Ala Gln Ala Ala 85 90 95
- Ala Gly Thr Val Asn Ile Gly Ala Ser Asp Ala Tyr Leu Ser Glu Gly 100 105 110
- Asp Met Ala Ala His Lys Gly Leu Met Asn Ile Ala Leu Ala Ile Ser 115 120 125
- Ala Gln Gln Val Asn Tyr Asn Leu Pro Gly Val Ser Glu His Leu Lys 130 135 140
- Leu Asn Gly Lys Val Leu Ala Ala Met Tyr Gln Gly Thr Ile Lys Thr 145 150 155 160
- Trp Asp Asp Pro Gln Ile Ala Ala Leu Asn Pro Gly Val Asn Leu Pro 165 170 175
- Gly Thr Ala Val Val Pro Leu His Arg Ser Asp Gly Ser Gly Asp Thr 180 . 185 190
- Phe Leu Phe Thr Gln Tyr Leu Ser Lys Gln Asp Pro Glu Gly Trp Gly 195 200 205
- Lys Ser Pro Gly Phe Gly Thr Thr Val Asp Phe Pro Ala Val Pro Gly 210 215 220
- Ala Leu Gly Glu Asn Gly Asn Gly Gly Met Val Thr Gly Cys Ala Glu 225 230 235 240
- Thr Pro Gly Cys Val Ala Tyr Ile Gly Ile Ser Phe Leu Asp Gln Ala 245 250 255
- Ser Gln Arg Gly Leu Gly Glu Ala Gln Leu Gly Asn Ser Ser Gly Asn 260 265 270

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166

365

Phe	Leu	Leu 275	Pro	Asp	Ala	Gln	Ser 280	Ile	Gln	Ala	Ala	Ala 285		Gly	Phe
Ala	Ser 290	Lys	Thr	Pro	Ala	Asn 295	Gln	Ala	Ile	Ser	Met 300	Ile	Asp	Gly	Pro
Ala 305	Pro	Asp	Gly	Tyr	Pro 310	Ile	Ile	Asn	Tyr	Glu 315	Tyr	Ala	Ile	Val	Asn 320
Asn	Arg	Gln	Lys	Asp 325	Ala	Ala	Thr	Ala	Gln 330	Thr	Leu	Gln	Ala	Phe 335	Leu
His	Trp	Ala	Ile 340	Thr	Asp	Gly	Asn	Lys 345	Ala	Ser	Phe	Leu	Asp 350	Gln	Val
His	Phe	Gln	Pro	Leu	Pro	Pro	Ala	Val	Val	Lys	Leu	Ser	Asp	Ala	Leu

360

Ile Ala Thr Ile Ser Ser 370

(2) INFORMATION FOR SEQ ID NO:149:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1993 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:149:

TGTTCTTCGA	CGGCAGGCTG	GTGGAGGAAG	GGCCCACCGA	ACAGCTGTTC	TCCTCGCCGA	60
AGCATGCGGA	AACCGCCCGA	TACGTCGCCG	GACTGTCGGG	GGACGTCAAG	GACGCCAAGC	120
GCGGAAATTG	AAGAGCACAG	AAAGGTATGG	CGTGAAAATT	CGTTTGCATA	CGCTGTTGGC	180
CGTGTTGACC	GCTGCGCCGC	TGCTGCTAGC	AGCGGCGGGC	TGTGGCTCGA	AACCACCGAG	240
CGGTTCGCCT	GAAACGGGCG	CCGGCGCCGG	TACTGTCGCG	ACTACCCCG	CGTCGTCGCC	300
GGTGACGTTG	GCGGAGACCG	GTAGCACGCT	GCTCTACCCG	CTGTTCAACC	TGTGGGGTCC	360
GGCCTTTCAC	GAGAGGTATC	CGAACGTCAC	GATCACCGCT	CAGGGCACCG	GTTCTGGTGC	420
CGGGATCGCG	CAGGCCGCCG	CCGGGACGGT	CAACATTGGG	GCCTCCGACG	CCTATCTGTC	480
GGAAGGTGAT	ATGGCCGCGC .	ACAAGGGGCT	GATGAACATC	GCGCTAGCCA	TCTCCGCTCA	540
GCAGGTCAAC	TACAACCTGC	CCGGAGTGAG	CGAGCACCTC	AAGCTGAACG	GAAAAGTCCT	600

GGCGGCCATG	TACCAGGGCA	CCATCAAAAC	CTGGGACGAC	CCGCAGATCG	CTGCGCTCAA	660
CCCCGGCGTG	AACCTGCCCG	GCACCGCGGT	AGTTCCGCTG	CACCGCTCCG	ACGGGTCCGG	720
TGACACCTTC	TTGTTCACCC	AGTACCTGTC	CAAGCAAGAT	CCCGAGGGCT	GGGGCAAGTC	780
GCCCGGCTTC	GGCACCACCG	TCGACTTCCC	GGCGGTGCCG	GGTGCGCTGG	GTGAGAACGG	840
CAACGGCGGC	ATGGTGACCG	GTTGCGCCGA	GACACCGGGC	TGCGTGGCCT	ATATCGGCAT	900
CAGCTTCCTC	GACCAGGCCA	GTCAACGGGG	ACTCGGCGAG	GCCCAACTAG	GCAATAGCTC	960
TGGCAATTTC	TTGTTGCCCG	ACGCGCAAAG	CATTCAGGCC	GCGGCGGCTG	GCTTCGCATC	1020
GAAAACCCCG	GCGAACCAGG	CGATTTCGAT	GATCGACGGG	CCCGCCCCGG	ACGGCTACCC	1080
GATCATCAAC	TACGAGTACG	CCATCGTCAA	CAACCGGCAA	AAGGACGCCG	CCACCGCGCA	1140
GACCTTGCAG	GCATTTCTGC	ACTGGGCGAT	CACCGACGGC	AACAAGGCCT	CGTTCCTCGA	1200
CCAGGTTCAT	TTCCAGCCGC	TGCCGCCCGC	GGTGGTGAAG	TTGTCTGACG	CGTTGATCGC	1260
GACGATTTCC	AGCTAGCCTC	GTTGACCACC	ACGCGACAGC	AACCTCCGTC	GGGCCATCGG	1320
GCTGCTTTGC	GGAGCATGCT	GGCCCGTGCC	GGTGAAGTCG	GCCGCGCTGG	CCCGGCCATC	1380
CGGTGGTTGG	GTGGGATAGG	TGCGGTGATC	CCGCTGCTTG	CGCTGGTCTT	GGTGCTGGTG	1440
GTGCTGGTCA	TCGAGGCGAT	GGGTGCGATC	AGGCTCAACG	GGTTGCATTT	CTTCACCGCC	1500
ACCGAATGGA	ATCCAGGCAA	CACCTACGGC	GAAACCGTTG	TCACCGACGC	GTCGCCCATC	1560
CGGTCGGCGC	CTACTACGGG	GCGTTGCCGC	TGATCGTCGG	GACGCTGGCG	ACCTCGGCAA	1620
TCGCCCTGAT	CATCGCGGTG	CCGGTCTCTG	TAGGAGCGGC	GCTGGTGATC	GTGGAACGGC	1680
TGCCGAAACG	GTTGGCCGAG	GCTGTGGGAA	TAGTCCTGGA	ATTGCTCGCC	GGAATCCCCA	1740
GCGTGGTCGT	CGGTTTGTGG	GGGGCAATGA	CGTTCGGGCC	GTTCATCGCT	CATCACATCG	1800
CTCCGGTGAT	CGCTCACAAC	GCTCCCGATG	TGCCGGTGCT	GAACTACTTG	CGCGGCGACC	1860
CGGGCAACGG	GGAGGGCATG	TTGGTGTCCG	GTCTGGTGTT	GGCGGTGATG	GTCGTTCCCA	1920
TTATCGCCAC	CACCACTCAT	GACCTGTTCC	GGCAGGTGCC	GGTGTTGCCC	CGGGAGGGCG	1980
CGATCGGGAA	TTC					1993

(2) INFORMATION FOR SEQ ID NO:150:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 374 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:150:

Met Lys Ile Arg Leu His Thr Leu Leu Ala Val Leu Thr Ala Ala Pro 1 5 10 15

Leu Leu Leu Ala Ala Gly Cys Gly Ser Lys Pro Pro Ser Gly Ser 20 25 30

Pro Glu Thr Gly Ala Gly Ala Gly Thr Val Ala Thr Thr Pro Ala Ser 35 40 45

Ser Pro Val Thr Leu Ala Glu Thr Gly Ser Thr Leu Leu Tyr Pro Leu 50 55 60 ;

Phe Asn Leu Trp Gly Pro Ala Phe His Glu Arg Tyr Pro Asn Vail Thr 65 70 75 80

Ile Thr Ala Gln Gly Thr Gly Ser Gly Ala Gly Ile Ala Gln Ala Ala 85 90 95

Ala Gly Thr Val Asn Ile Gly Ala Ser Asp Ala Tyr Leu Ser Glu Gly
100 105 110

Asp Met Ala Ala His Lys Gly Leu Met Asn Ile Ala Leu Ala Ile Ser 115 120 125

Ala Gln Gln Val Asn Tyr Asn Leu Pro Gly Val Ser Glu His Leu Lys 130 135 140

Leu Asn Gly Lys Val Leu Ala Ala Met Tyr Gln Gly Thr Ile Lys Thr 145 150 155 160

Trp Asp Asp Pro Gln Ile Ala Ala Leu Asn Pro Gly Val Asn Leu Pro 165 170 175

Gly Thr Ala Val Val Pro Leu His Arg Ser Asp Gly Ser Gly Asp Thr 180 185 190

Phe Leu Phe Thr Gln Tyr Leu Ser Lys Gln Asp Pro Glu Gly Trp Gly 195 200 205

Lys Ser Pro Gly Phe Gly Thr Thr Val Asp Phe Pro Ala Val Pro Gly 210 215 220

Ala Leu Gly Glu Asn Gly Asn Gly Gly Met Val Thr Gly Cys Ala Glu 225 230 235 240

Thr Pro Gly Cys Val Ala Tyr Ile Gly Ile Ser Phe Leu Asp Gln Ala 245 250 255

Ser Gln Arg Gly Leu Gly Glu Ala Gln Leu Gly Asn Ser Ser Gly Asn 265 Phe Leu Leu Pro Asp Ala Gln Ser Ile Gln Ala Ala Ala Gly Phe 280 Ala Ser Lys Thr Pro Ala Asn Gln Ala Ile Ser Met Ile Asp Gly Pro 300 295 290 Ala Pro Asp Gly Tyr Pro Ile Ile Asn Tyr Glu Tyr Ala Ile Val Asn 315 310 Asn Arg Gln Lys Asp Ala Ala Thr Ala Gln Thr Leu Gln Ala Phe Leu 330 325 His Trp Ala Ile Thr Asp Gly Asn Lys Ala Ser Phe Leu Asp Gln Val 345 His Phe Gln Pro Leu Pro Pro Ala Val Val Lys Leu Ser Asp Ala Leu 360

Ile Ala Thr Ile Ser Ser 370

(2) INFORMATION FOR SEQ ID NO:151:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1777 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:151:

GGTCTTGACC	ACCACCTGGG	TGTCGAAGTC	GGTGCCCGGA	TTGAAGTCCA	GGTACTCGTG	60
GGTGGGGCGG	GCGAAACAAT	AGCGACAAGC	ATGCGAGCAG	CCGCGGTAGC	CGTTGACGGT	120
GTAGCGAAAC	GGCAACGCGG	CCGCGTTGGG	CACCTTGTTC	AGCGCTGATT	TGCACAACAC	180
CTCGTGGAAG	GTGATGCCGT	CGAATTGTGG	CGCGCGAACG	CTGCGGACCA	GGCCGATCCG	240
CTGCAACCCG	GCAGCGCCCG	TCGTCAACGG	GCATCCCGTT	CACCGCGACG	GCTTGCCGGG	300
CCCAACGCAT	ACCATTATTC	GAACAACCGT	TCTATACTTT	GTCAACGCTG	GCCGCTACCG	360
AGCGCCGCAC	AGGATGTGAT	ATGCCATCTC	TGCCCGCACA	GACAGGAGCC	AGGCCTTATG	420
ACAGCATTCG	GCGTCGAGCC	CTACGGGCAG	CCGAAGTACC	TAGAAATCGC	CGGGAAGCGC	480
ATGGCGTATA	TCGACGAAGG	CAAGGGTGAC	GCCATCGTCT	TTCAGCACGG	CAACCCCACG	540

AGCTATGGCG AGCAACGAGA CTTTTTGTTC GCGCTCTGGG ATGCGCTCGA CCTCGGCGAC CACGTGGTAC TGGTGCTGCA CGACTGGGGC TCGGCGCTCG GCTTCGACTG GGCTAACCAG CATCGCGACC GAGTGCAGGG GATCGCGTTC ATGGAAGCGA TCGTCACCCC GATGACGTGG GCGGACTGGC CGCCGGCCGT GCGGGGTGT TTCCAGGGTT TCCGATCGCC TCAAGGCGAG CCAATGGCGT TGGAGCACAA CATCTTTGTC GAACGGGTC TGCCCGGGGC GATCCTGCGA CAGCTCAGCG ACGAGGAAAT GAACCACTAT CGGCGGCCAT TCGTGAACGG CGGCGAGGAC CGTCGCCCCA CGTTGTCGTG GCCACGAAAC CTTCCAATCG ACGGTGAGCC CGCCGAGGTC GTCGCGTTGG TCAACGAGTA CCGGAGCTGG CTCGAGGAAA CCGACATGCC GAAACTGTTC ATCAACGCCG AGCCCGGCG GATCATCACC GGCCGCATCC GTGACTATGT CAGGAGCTGG CCCAACCAGA CCGAAATCAC AGTGCCCGGC GTGCATTTCG TTCAGGAGGA CAGCGATGGC GTCGTATCGT GGGCGGGCGC TCGGCAGCAT CGGCGACCTG GGAGCGCTTC CATTTCACGA GACCAAGAAT GTGATTTCCG GCGAAGGCGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT GGCTCCGGCA GAGATTCAC AGGACAAAGG CACCAATCGC AGCCGTTCC TTCGCAACGA GGCTCGGCAAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCC TTCGCAACGA TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG ATCATCCCAC AGGGCTGCT TCACAACGCC CACATGACT ACGTGTTACT AGCGCCCAGC CCCGATCCGC AGGGGTGCT TCACAACGCC CACATGACT ACGTGTTACT AGCGCCCAGC CCCGATCCGG AGGAGGTGCT CTCACAACGCC CACATGACT ACGTGTTACT AGCGCCCAGC CCCGATCCGG AGGAGGTGCT CTCACAACGCC CACATGACT ACGTGTTACT AGCGCCCAGC CCCGATCCGG AGGAGGTGCT CTCACAACGCC CACATGACT ACGTGTTACT AGCGCCCAGC CCCGATCCGG AGGAGGTGCT CGCCGCCCCC CCCCCCCCCC	CGTCTTA	CT TGTG	GCGCAA	CATCATGCC	G CACTTGGAAC	GGCTGGGCCG	GCTGGTGGCC	600
CACGTGGTAC TGGTGCTGCA CGACTGGGGC TCGGCGCTCG GCTTCGACTG GGCTAACCAG CATCGCGACC GAGTGCAGGG GATCGCGTTC ATGGAAGCGA TCGTCACCCC GATGACGTGG GCGGACTGGC CGCCGGCCGT GCGGGGTGTG TTCCAGGGTT TCCGATCGCC TCAAGGCGAG GCGGACTGGC CGCCGGCCGT GCGGGGTGTG TTCCAGGGTT TCCGATCGCC TCAAGGCGAG CCAATGGCGT TGGAGCACAA CATCTTTGTC GAACGGGTGC TGCCCGGGGC GATCCTGCGA CAGCTCAGCG ACGAGGAAAT GAACCACTAT CGGCGGCCAT TCGTGAACGG CGGCGAGGAC CGTCGCCCCA CGTTGTCGTG GCCACGAAAC CTTCCAATCG ACGGTGAGCC CGCCGAGGTC GTCGCGTTGG TCAACGAGTA CCGGAGCTGG CTCGAGGAAA CCGACATGCC GAAACTGTTC ATCAACGCCG AGCCCGGCGC GATCATCACC GGCCGATCC GTGACTATGT CAGGAGCTGG CCCAACCAGA CCGAAATCAC AGTGCCCGGC GTGCATTTCG TTCAGGAGGA CAGCGATGGC GTCGTATCGT GGGCGGGCGC TCGGCAGCAT CGGCGACCTG GGAGCGCTCT CATTTCACGA GACCAAGAAT GTGATTTCCG GCGAAGGGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT GGCTCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG ATCATCCCAC AGGGCTGCT TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC ATCATCCCAC AGGGCTGCT TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC CCCGATCCGG AGGAGGTGCT GCCGCCCCC CACATGTACT ACGTGTTACT AGCGCCCAGC TTACCTCCAC AGGGCTGCT CTCACAACCCC CACATGTACT ACGTGTTACT AGCGCCCAGC CCCGATCCGG AGGAGGTGCT GCCGCCCCCC CACATGTACT ACGTGTTACT AGCGCCCAGC TTACCTCCGC AGGAGGTGCT CCCCCCCCC CACCTCCCTTTCCTTT	GCGATCT	GA TCGG	GATGGG	CGCGTCGGA	C AAGCTCAGCC	CATCGGGACC	CGACCGCTAT	660
CATCGCGACC GAGTGCAGGG GATCGCGTTC ATGGAAGCGA TCGTCACCCC GATGACGTGG GCGGACTGGC CGCCGGCCGT GCGGGGTTG TTCCAGGGTT TCCGATCGCC TCAAGGCGAG GCGGACTGGC CGCCGGCCGT GCGGGGTTG TTCCAGGTT TCCGATCGCC TCAAGGCGAG CCAATGGCGT TGGAGCACAA CATCTTTGTC GAACGGGTGC TGCCCGGGGC GATCCTGCGA GCGACTCAGCG ACGAGGAAAT GAACCACTAT CGGCGGCCAT TCGTGAACGG CGGCGAGGAC CGTCGCCCCA CGTTGTCGTG GCCACGAAAC CTTCCAATCG ACGGTGAGCC CGCCGAGGTC GTCGCGTTGG TCAACGAGTA CCGGAGCTGG CTCGAGGAAA CCGACATGCC GAAACTGTTC ATCAACGCCG AGCCCGGCCG GATCATCACC GGCCGCATCC GTGACTATGT CAGGAGCTGG CCCAACCAGA CCGAAATCAC AGTGCCCGGC GTGCATTTCG TTCAGGAGGA CAGCGATGGC GTCGTATCGT GGGCGGGCGC TCGGCAGCAT CGGCGACCTG GGAGCGCTCT CATTTCACGA GACCAAGAAT GTGATTTCCG GCGAAGGCGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT GCTCCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT TGGAACCAAC ACGCGCTCC CGGCGGACCC GAGCGGTCC GCGCCGTCCC GCCTTCCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG ATCATCCCAC AGGGCTGCT CTCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC ATCATCCCAC AGGGCTGCT CTCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC CCCGATCCGG AGGAGGTGCT GCCGCCCCC CACATGTACT ACGTGTTACT AGCGCCCAGC TTACGTGCCGC TTCACGATTC GCCGGCCCCC CCTCCCTT TCACGTCCCCT TCACCAATCC CGCCCCCCC CACATCCCTT TACGTGCCCC TTCACCATTC GCCGCCCCC CCTCCCTT TCACGTCCCTC TTCACCATTC GCCGCCCCC CCTCCCTTT TACGTGCCCC TTCACCATTC GCCGCCCCC CCTCCCTTT TACGTGCCCC TTCACCATTC GCCGCCCCC CCTCCCTTT TCACGTCCCCC TTCACCATTC GCCGCCCCC CCTCCCCTTT TACGTGCCCC TTCACCATTC GCCGCCCCC CCTCCCTTT TACGTGCCCC TTCACCATTC GCCGCCCCC CCTCCCTTT TACGTGCCCC TTCACCATTC GCCGCCCCC CCTCCCTTT TACGTGCCCC TTCACCATTC ACGACCAACGC GGTCTTTCAT TACGTGCCCCC TTCACCATTC ACGACCAACGC GGTCTTTCAT TACGTCCCCC TTCACCATTC GCCGCCCCC CCCCCCCCCC	GCTATGG	CG AGCA	ACGAGA	CTTTTTGTTC	C GCGCTCTGGG	ATGCGCTCGA	CCTCGGCGAC	720
GCGGACTGGC CGCCGGCCGT GCGGGGTGTG TTCCAGGGTT TCCGATCGCC TCAAGGCGAG CCAATGGCGT TGGAGCACAA CATCTTTGTC GAACGGGTGC TGCCCGGGGC GATCCTGCGA CAGCTCAGCG ACGAGGAAAT GAACCACTAT CGGCGGCCAT TCGTGAACGG CGGCGAGGAC CGTCGCCCCA CGTTGTCGTG GCCACGAAAC CTTCCAATCG ACGGTGAGCC CGCCGAGGTC GTCGCGTTGG TCAACGAGTA CCGGAGCTGG CTCGAGGAAA CCGACATGCC GAAACTGTTC ATCAACGCCG AGCCCGGCG GATCATCACC GGCCGCATCC GTGACTATGT CAGGAGCTGG CCCAACCAGA CCGAAATCAC AGTGCCCGGC GTGCATTTCG TTCAGGAGGA CAGCGATGGC GTCGTATCGT GGGCGGGCG TCGGCAGCAT CGGCGACCTG GGAGCGCTCT CATTTCACGA GACCAAGAAT GTGATTTCCG GCGAAGAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA GCTCCGGGCA GAGATCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA GGTCGACAAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG ATCATCCCAC AGGGCTGCT TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC ATCATCCCAC AGGGCTGCT TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC CCCGATCGGG AGGAGGTGCT GCCGGCCCCCC CCTCCCTT TACGTGCCGC TTCACGATTC GCCGGCCCCCC CCTCCCTTT TACGTGCCCGC TTCACGATTC GCCGGCCCCCC CCTCCCTTT TACGTGCCCGC TTCACGATTC GCCGGCCCCCC CCTCCCTTT TACGTGCCCGC TTCACGATTC GCCGGCCCCCC CCTCCCTTT TACGTGCCCGC TTCACGATTC GCCGGCCCCCC CCTCCCTTT TACGTGCCCCC TTCACGATTC GCCGGCCCCCC CCTCCCTTT TACGTGCCCCC TTCACGATTC GCCGGCCCCCC CCTCCCTTT TACGTGCCCCC TTCACGATTC GCCGCCCCCC CCTCCCTTT TACGTGCCCCC TTCACGATTC GCCGCCCCCC CCTCCCTTT TACGTGCCCCC TTCACGATTC GCCGCCCCCC CCTCCCCTTT TACGTGCCCCC TTCACGATTC GCCGCCCCCC CCTCCCTTT TACGTGCCCCC TTCACGATTC GCCGCCCCCCC CCTCCCCTTT TACGTGCCCCC TTCACGATTC GCCCCCCCCC CCTCCCCTTT TACGTGCCCCC TTCACGATTC GCCCCCCCCCCCCCCCC	ACGTGGT	AC TGGT	GCTGCA	CGACTGGGG	TCGGCGCTCG	GCTTCGACTG	GGCTAACCAG	780
CCAATGGCGT TGGAGCACAA CATCTTTGTC GAACGGGTGC TGCCCGGGGC GATCCTGCGA CAGCTCAGCG ACGAGGAAAT GAACCACTAT CGGCGGCCAT TCGTGAACGG CGGCGAGGAC CGTCGCCCCA CGTTGTCGTG GCCACGAAAC CTTCCAATCG ACGGTGAGCC CGCCGAGGTC GTCGCGTTGG TCAACGAGTA CCGGAGCTGG CTCGAGGAAA CCGACATGCC GAAACTGTTC ATCAACGCCG AGCCCGGCGC GATCATCACC GGCCGCATCC GTGACTATGT CAGGAGCTGG CCCCAACCAGA CCGAAATCAC AGTGCCCGGC GTGCATTTCG TTCAGGAGGA CAGCGATGGC GTCGTATCGT GGGCGGGCGC TCGGCAGCAT CGGCGACCTG GGAGCGCTCT CATTTCACGA GACCAAGAAT GTGATTTCCG GCGAAGGCGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT 136 GCTCCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC GCCGATCGGG AGGAGGTGCT GCCGGCGCCC CCTCCCCTC TACCGTGCCGC TTCACCACTC GCCGCCCCC CCTCCCCTC TACCGTGCCGC TTCACCACTTC GCCGCCCCC CCTCCCCTC TACCGTGCCGC TTCACCACTTC GCCGCCCCC CCTCCCCTC TACCGTGCCGC TTCACCACTTC GCCGCCCCC CCTCCCCTC TACCGTGCCGC TTCACCACTTC GCCGCCCCCC CCTCCCCTCC	ATCGCGA	CC GAGT	GCAGGG	GATCGCGTTC	ATGGAAGCGA	TCGTCACCCC	GATGACGTGG	840
CAGCTCAGCG ACGAGGAAAT GAACCACTAT CGGCGGCCAT TCGTGAACGG CGGCGAGGAC CGTCGCCCCA CGTTGTCGTG GCCACGAAAC CTTCCAATCG ACGGTGAGCC CGCCGAGGTC GTCGCGTTGG TCAACGAGTA CCGGAGCTGG CTCGAGGAAA CCGACATGCC GAAACTGTTC ATCAACGCCG AGCCCGGCGC GATCATCACC GGCCGCATCC GTGACTATGT CAGGAGCTGG CCCAACCAGA CCGAAATCAC AGTGCCCGGC GTGCATTTCG TTCAGGAGGA CAGCGATGGC GTCGTATCGT GGGCGGGCGC TCGGCAGCAT CGGCGACCTG GGAGCGCTCT CATTTCACGA GACCAAGAAT GTGATTTCCG GCGAAGGCGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT GCTCCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT GCCTTTCTAT GGGCTCAGTT CGAGGAAACG GAGCGATCA CGCGTTCCC ATTGGACCTA TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC GCCGATCGGG AGGAGGTGCT GGCCGCCCC CCTCCCTT TACGTGCCGC TTCACGATTC GCCGCCCCC CCTCCCTTT TACGTGCCCC TTCACGATTC GCCGCCCCC CCTCCCCTTT TACGTGCCCC TTCACGATTC GCCGCCCCC CCTCCCCTTT TACGTGCCCCC TTCACGATTC GCCGCCCCC CCTCCCCTTT TACGTGCCCCC TTCACGATTC GCCCCCCC CCTCCCCTTT TACGTGCCCCC TTCACGATTC GCCCCCCC CCTCCCCTTT TACGTGCCCCC TTCACGATTC GCCCCCC CCTCCCCTTT TACGTGCCCCC TTCACGATTC GCCCCCC CCTCCCCTTT TACGTGCCCC TTCACGATTC GCCCCCCC CCTCCCCTTT TACGTGCCCCC TTCACGATTC GCCCCCC CCTCCCCTTT TACGTGCCCCC TTCACGATTC GCCCCCCC CCTCCCCTTT TACGTGCCCCC TTCACCCCCC CCCCCCCCC CCTCCCCCCCCCC	CGGACTG	C CGCC	GGCCGT	GCGGGGTGTG	TTCCAGGGTT	TCCGATCGCC	TCAAGGCGAG	900
CGTCGCCCCA CGTTGTCGTG GCCACGAAAC CTTCCAATCG ACGGTGAGCC CGCCGAGGTC GTCGCGTTGG TCAACGAGTA CCGGAGCTGG CTCGAGGAAA CCGACATGCC GAAACTGTTC ATCAACGCCG AGCCCGGCGC GATCATCACC GGCCGCATCC GTGACTATGT CAGGAGCTGG CCCAACCAGA CCGAAATCAC AGTGCCCGGC GTGCATTTCG TTCAGGAGGA CAGCGATGGC GTCGTATCGT GGGCGGGCGC TCGGCAGCAT CGGCGACCTG GGAGCGCTCT CATTTCACGA GACCAAGAAT GTGATTTCCG GCGAAGGCGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT GCTCCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC TACGTGCCGC TTCACGATTC GCCGCCCCCC CCTCCCTT TACGTGCCGC TTCACGATTC GCCGCCCCCC CCTCCCTTT TACGTGCCCCCCCCC TCCACGATTC GCCGCCCCCCC CCTCCCTTT TACGTGCCCCCCCC TCCACGATTC GCCGCCCCCCC TCCTCCCTTT TACGTGCCCCCCCCC TCCACGATTC GCCGCCCCCCC TCCTCCCTTT TACGTGCCCCCCCCC TCCACGATTC GCCGCCCCCC TCCTCCCTTT TACGTGCCCCCCCCCC TCCACGATTC GCCGCCCCCCC TCCCCCTTT TACGTGCCCCCCCCCCCC TCCACCCCCCCC TCCCCCTTCCCCTTT TACGTGCCCCCCCCCCCCCCCCC	CAATGGC	T TGGA	GCACAA	CATCTTTGTC	GAACGGGTGC	TGCCCGGGGC	GATCCTGCGA	960
GTCGCGTTGG TCAACGAGTA CCGGAGCTGG CTCGAGGAAA CCGACATGCC GAAACTGTTC ATCAACGCCG AGCCCGGCGC GATCATCACC GGCCGCATCC GTGACTATGT CAGGAGCTGG CCCAACCAGA CCGAAATCAC AGTGCCCGGC GTGCATTTCG TTCAGGAGGA CAGCGATGGC GTCGTATCGT GGGCGGGCGC TCGGCAGCAT CGGCGACCTG GGAGCGCTCT CATTTCACGA GACCAAGAAT GTGATTTCCG GCGAAGGCGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT GCTCCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA TGGAAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC GCCGATCGGG AGGAGGTGCT GGCGCGCCC CCTCCCTT TACGTGCCGC TTCACGATTC GCCGGCCCCC GCTCCCTT TACGTGCCCCC TTCACGATTC GCCGGCCCCC GCTCCCTT TACGTGCCCCC TTCACGATTC GCCGGCCCCC GCTCCCTT TACGTGCCCCC TTCACGATTC GCCGGCCCCC GCTCCCTT TACGTGCCCCC TTCACGATTC GCCGCCCCC GCTCCCTT TACGTGCCCCC TTCACGATTC GCCGCCCCC GCTCCCTT TACGTGCCCCC TTCACGATTC GCCGCCCCC GCTCCCTT TACGTGCCCCC TTCACGATTC GCCGCCCCC GCTCCCTT TACGTGCCCCC TTCACGATTC GCCGCCCCCC GCTCCCTT TACGTGCCCCCCCC TTCACGATTC GCCGCCCCCC GCTCCCTT TACGTGCCCCCC TTCACGATTC GCCGCCCCC GCTCCCTT TACGTGCCCCCCCC TTCACGATTC GCCCCCCCC GCTCCCTT TACGTGCCCCCCCCC TTCACGATTC GCCCCCCCCCCCC	AGCTCAGO	G ACGA	GGAAAT	GAACCACTAT	CGGCGGCCAT	TCGTGAACGG	CGGCGAGGAC	1020
ATCAACGCCG AGCCCGGCGC GATCATCACC GGCCGCATCC GTGACTATGT CAGGAGCTGG 120 CCCAACCAGA CCGAAATCAC AGTGCCCGGC GTGCATTTCG TTCAGGAGGA CAGCGATGGC 120 GTCGTATCGT GGGCGGGCGC TCGGCAGCAT CGGCGACCTG GGAGCGCTCT CATTTCACGA 132 GACCAAGAAT GTGATTTCCG GCGAAGGCGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT 138 GCTCCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA 144 GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT 150 GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA 156 TTGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG 162 ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC 166 GCCGATCGGG AGGAGGTGCT GGCGCCCCC CACATGTACT ACGTGTTACT AGCGCCCAGC 166 TTCAGGAGTCCC TTCACGATTC GCCGGCCCCC CGTCCCGTT	GTCGCCC	A CGTT	STCGTG	GCCACGAAAC	CTTCCAATCG	ACGGTGAGCC	CGCCGAGGTC	1080
CCCAACCAGA CCGAAATCAC AGTGCCCGGC GTGCATTTCG TTCAGGAGGA CAGCGATGGC 126 GTCGTATCGT GGGCGGCGC TCGGCAGCAT CGGCGACCTG GGAGCGCTCT CATTTCACGA 132 GACCAAGAAT GTGATTTCCG GCGAAGGCGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT 136 GCTCCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA 144 GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT 156 GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA 156 TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG 162 ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC 168 GCCGATCGGG AGGAGGTGCT GGCGCGCCC CCCCGCGTCCG TTACGTGCCGC TTCACGATTC GCCGCCCCC CCTCCGTT	rcgcgtte	G TCAA	CGAGTA	CCGGAGCTGG	CTCGAGGAAA	CCGACATGCC	GAAACTGTTC	1140
GTCGTATCGT GGGCGGCGC TCGGCAGCAT CGGCGACCTG GGAGCGCTCT CATTTCACGA GACCAAGAAT GTGATTTCCG GCGAAGGCGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT GCTCCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC GCCGATCGGG AGGAGGTGCT GGCGCCCCC CCTCGCTT TACGTGCCGC TTCACGATTC GCCGCCCCC CCTCGCTT TACGTGCCGC TTCACGATTC GCCGCCCCC CCTCGCTT	I'CAACGCC	G AGCCC	CGGCGC	GATCATCACC	GGCCGCATCC	GTGACTATGT	CAGGAGCTGG	1200
GACCAAGAAT GTGATTTCCG GCGAAGGCGG CGCCCTGCTT GTCAACTCAT AAGACTTCCT 138 GCTCCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA 144 GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT 156 GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA 156 TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG 162 ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC 168 GCCGATCGGG AGGAGGTGCT GGCGCCCGC CACAGGCGAAG GTATAGGCGC GGTCTTTCAT 174 TACGTGCCGC TTCACGATTC GCCGCCCGC CCTCCGT	CCAACCAG	A CCGA	ATCAC	AGTGCCCGGC	GTGCATTTCG	TTCAGGAGGA	CAGCGATGGC	1260
GCTCCGGGCA GAGATTCTCA GGGAAAAGGG CACCAATCGC AGCCGCTTCC TTCGCAACGA 144 GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT 150 GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA 156 TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG 162 ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC 168 GCCGATCGGG AGGAGGTGCT GGCGCCCCC CACAGGCGAAG GTATAGGCGC GGTCTTTCAT 174 TACGTGCCGC TTCACGATTC GCCGGCCCCC CCTCGCT	CGTATCG	T GGGCG	GGCGC	TCGGCAGCAT	CGGCGACCTG	GGAGCGCTCT	CATTTCACGA	1320
GGTCGACAAA TATACGTGGC AGGACAAAGG TCTTCCTATT TGCCCAGCGA ATTAGTCGCT 150 GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA 156 TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG 162 ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC 168 GCCGATCGGG AGGAGGTGCT GGCGCCCCC CCTCGCTT TACGTGCCGC TTCACGATTC GCCGCCCCCC CCTCGCTT	ACCAAGAA	T GTGAT	TTCCG	GCGAAGGCGG	CGCCCTGCTT	GTCAACTCAT	AAGACTTCCT	1380
GCCTTTCTAT GGGCTCAGTT CGAGGAAGCC GAGCGGATCA CGCGTATCCG ATTGGACCTA 156 TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG 162 ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC 168 GCCGATCGGG AGGAGGTGCT GGCGCGCCCG CCTCGCTT TACGTGCCGC TTCACGATTC GCCGGCCCGC CCTCGCTT	TCCGGGC	A GAGAT	TCTCA	GGGAAAAGGG	CACCAATCGC	AGCCGCTTCC	TTCGCAACGA	1440
TGGAACCGGT ATCATGAAAG CTTCGAATCA TTGGAACAGC GGGGGCTCCT GCGCCGTCCG 162 ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC 168 GCCGATCGGG AGGAGGTGCT GGCGCGCCG ACGAGCGAAG GTATAGGCGC GGTCTTTCAT 174 TACGTGCCGC TTCACGATTC GCCGCCCGC CCTGCGT	TCGACAA	A TATAC	GTGGC A	AGGACAAAGG	TCTTCCTATT	TGCCCAGCGA	ATTAGTCGCT	1500
ATCATCCCAC AGGGCTGCTC TCACAACGCC CACATGTACT ACGTGTTACT AGCGCCCAGC 168 GCCGATCGGG AGGAGGTGCT GGCGCGCTCTG ACGAGCGAAG GTATAGGCGC GGTCTTTCAT 174 TACGTGCCGC TTCACGATTC GCCGCCCCC CCTCGCT	CTTTCTA	r GGGCT	CAGTT (CGAGGAAGCC	GAGCGGATCA	CGCGTATCCG	ATTGGACCTA	1560
GCCGATCGGG AGGAGGTGCT GGCGCGCCGC ACGAGCGAAG GTATAGGCGC GGTCTTTCAT TACGTGCCGC TTCACGATTC GCCGCCCGC CCTCGCT	GAACCGG'	T ATCAT	GAAAG (CTTCGAATCA	TTGGAACAGC	GGGGGCTCCT	GCGCCGTCCG	1620
TACGTGCCGC TTCACGATTC GCCGCCCCC CCTCCCT	CATCCCA	AGGGC'	TGCTC 1	rcacaacgcc	CACATGTACT	ACGTGTTACT	AGCGCCCAGC	1680
TACGTGCCGC TTCACGATTC GCCGGCCGGG CGTCGCT	CGATCGG	G AGGAG	GTGCT G	GCGCGTCTG	ACGAGCGAAG	GTATAGGCGC	GGTCTTTCAT	1740
177	CGTGCCG	TTCAC	GATTC G	CCGGCCGGG	CGTCGCT			1777

(2) INFORMATION FOR SEQ ID NO:152:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 324 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE	DESCRIPTION:	SEQ	ΙD	NO:152:
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GAGATTGAAT	CGTACCGGTC	TCCTTAGCGG	CTCCGTCCCG	TGAATGCCCA	TATCACGCAC	60
GGCCATGTTC	TGGCTGTCGA	CCTTCGCCCC	ATGCCCGGAC	GTTGGTAAAC	CCAGGGTTTG	120
ATCAGTAATT	CCGGGGGACG	GTTGCGGGAA	GGCGGCCAGG	ATGTGCGTGA	GCCGCGGCGC	180
CGCCGTCGCC	CAGGCGACCG	CTGGATGCTC	AGCCCCGGTG	CGGCGACGTA	GCCAGCGTTT	240
GGCGCGTGTC	GTCCACAGTG	GTACTCCGGT	GACGACGCGG	CGCGGTGCCT	GGGTGAAGAC	300
CGTGACCGAC	GCCGCCGATT	CAGA				324

(2) INFORMATION FOR SEQ ID NO:153:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1338 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:153:

GCGGTACCGC CGCGTTGCGC TG	GCACGGGA	CCTGTACGAC	CTGAACCACT	TCGCCTCGCG	60
AACGATTGAC GAACCGCTCG TG	CGGCGGCT	GTGGGTGCTC	AAGGTGTGGG	GTGATGTCGT	120
CGATGACCGG CGCGGCACCC GG	CCACTACG	CGTCGAAGAC	GTCCTCGCCG	CCCGCAGCGA	180
GCACGACTTC CAGCCCGACT CG	ATCGGCGT	GCTGACCCGT	CCTGTCGCTA	TGGCTGCCTG	240
GGAAGCTCGC GTTCGGAAGC GA	TTTGCGTT	CCTCACTGAC	CTCGACGCCG	ACGAGCAGCG	300
GTGGGCCGCC TGCGACGAAC GG	CACCGCCG	CGAAGTGGAG	AACGCGCTGG	CGGTGCTGCG	360
GTCCTGATCA ACCTGCCGGC GA	TCGTGCCG	TTCCGCTGGC	ACGGTTGCGG	CTGGACGCGG	420
CTGAATCGAC TAGATGAGAG CA	GTTGGGCA	CGAATCCGGC	TGTGGTGGTG	AGCAAGACAC	480
GAGTACTGTC ATCACTATTG GA	TGCACTGG	ATGACCGGCC	TGATTCAGCA	GGACCAATGG	540
AACTGCCCGG GGCAAAACGT CT	CGGAGATG	ATCGGCGTCC	CCTCGGAACC	CTGCGGTGCT	600
GGCGTCATTC GGACATCGGT CC	GGCTCGCG	GGATCGTGGT	GACGCCAGCG	CTGAAGGAGT	660
GGAGCGCGGC GGTGCACGCG CT	GCTGGACG	GCCGGCAGAC	GGTGCTGCTG	CGTAAGGGCG	720
GGATCGGCGA GAAGCGCTTC GA	GGTGGCGG	CCCACGAGTT	CTTGTTGTTC	CCGACGGTCG	780
CGCACAGCCA CGCCGAGCGG GT	TCGCCCCG	AGCACCGCGA	CCTGCTGGGC	ccecceccc	840

CCGACAGCAC	CGACGAGTGT	GTGCTACTGC	GGGCCGCAGC	GAAAGTTGTT	GCCGCACTGC	900
CGGTTAACCG	GCCAGAGGGT	CTGGACGCCA	TCGAGGATCT	GCACATCTGG	ACCGCCGAGT	960
CGGTGCGCGC	CGACCGGCTC	GACTTTCGGC	CCAAGCACAA	ACTGGCCGTC	TTGGTGGTCT	1020
CGGCGATCCC	GCTGGCCGAG	CCGGTCCGGC	TGGCGCGTAG	GCCCGAGTAC	GGCGGTTGCA	1080
CCAGCTGGGT	GCAGCTGCCG	GTGACGCCGA	CGTTGGCGGC	GCCGGTGCAC	GACGAGGCCG	1140
CGCTGGCCGA	GGTCGCCGCC	CGGGTCCGCG	AGGCCGTGGG	TTGACTGGGC	GGCATCGCTT	1200
GGGTCTGAGC	TGTACGCCCA	GTCGGCGCTG	CGAGTGATCT	GCTGTCGGTT	CGGTCCCTGC	1260
TGGCGTCAAT	TGACGGCGCG	GGCAACAGCA	GCATTGGCGG	CGCCATCCTC	CGCGCGGCCG	1320
GCGCCCACCG	CTACAACC				f	1338

(2) INFORMATION FOR SEQ ID NO:154:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 321 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:154:

CCGGCGGCAC	CGGCGGCACC	GGCGGTACCG	GCGGCAACGG	CGCTGACGCC	GCTGCTGTGG	60
TGGGCTTCGG	CGCGAACGGC	GACCCTGGCT	TCGCTGGCGG	CAAAGGCGGT	AACGGCGGAA	120
TAGGTGGGGC	CGCGGTGACA	GGCGGGGTCG	CCGGCGACGG	CGGCACCGGC	GGCAAAGGTG	180
GCACCGGCGG	TGCCGGCGGC	GCCGGCAACG	ACGCCGGCAG	CACCGGCAAT	CCCGGCGGTA	240
AGGGCGGCGA	CGGCGGGATC	GGCGGTGCCG	GCGGGGCCGG	CGGCGCGGCC	GGCACCGGCA	300
ACGGCGGCCA	TGCCGGCAAC	С				321

(2) INFORMATION FOR SEQ ID NO:155:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 492 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:155:

GAAGACCCGG	CCCCGCCATA	TCGATCGGCT	CGCCGACTAC	TTTCGCCGAA	CGTGCACGCG	60
GCGGCGTCGG	GCTGATCATC	ACCGGTGGCT	ACGCGCCCAA	CCGCACCGGA	TGGCTGCTGC	120
CGTTCGCCTC	CGAACTCGTC	ACTTCGGCGC	AAGCCCGACG	GCACCGCCGA	ATCACCAGGG	180
CGGTCCACGA	TTCGGGTGCA	AAGATCCTGC	TGCAAATCCT	GCACGCCGGA	CGCTACGCCT	240
ACCACCCACT	TGCGGTCAGC	GCCTCGCCGA	TCAAGGCGCC	GATCACCCCG	TTTCGTCCGC	300
GAGCACTATC	GGCTCGCGGG	GTCGAAGCGA	CCATCGCGGA	TTTCGCCCGC	TGCGCGCAGT	360
TGGCCCGCGA	TGCCGGCTAC	GACGGCGTCG	AAATCATGGG	CAGCGAAGGG	TATCTGCTCA	420
ATCAGTTCCT	GGCGCCGCGC	ACCAACAAGC	GCACCGACTC	GTGGGGCGGC	ACACCGGCCA	480
ACCGTCGCCG	GT					492

(2) INFORMATION FOR SEQ ID NO:156:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 536 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:156:

Phe Ala Gln His Leu Val Glu Gly Asp Ala Val Glu Leu Trp Arg Ala
1 5 10 15

Asn Ala Ala Asp Gln Ala Asp Pro Leu Gln Pro Gly Ser Ala Arg Arg 20 25 30

Gln Arg Ala Ser Arg Ser Pro Arg Arg Leu Ala Gly Pro Asn Ala Tyr 35 40 45

His Tyr Ser Asn Asn Arg Ser Ile Leu Cys Gln Arg Trp Pro Leu Pro 50 55 60

Ser Ala Ala Gln Asp Val Ile Cys His Leu Cys Pro His Arg Gln Glu 65 70 75 80

Pro Gly Leu Met Thr Ala Phe Gly Val Glu Pro Tyr Gly Gln Pro Lys 85 90 95

Tyr Leu Glu Ile Ala Gly Lys Arg Met Ala Tyr Ile Asp Glu Gly Lys

			100					105					110		
Gly	Asp	Ala 115	Ile	Val	Phe	Gln	Ніs 120	Gly	Asn	Pro	Thr	Ser 125	Ser	Tyr	Leu
Trp	Arg 130	Asn	Ile	Met	Pro	His 135	Leu	Glu	Gly	Leu	Gly 140	Arg	Leu	Val	Ala
Cys 145	Asp	Leu	Ile	Gly	Met 150	Gly	Ala	Ser	Asp	Lys 155	Leu	Ser	Pro	Ser	Gly 160
Pro	Asp	Arg	Tyr	Ser 165	Tyr	Gly	Glu	Gln	Arg 170	Asp	Phe	Leu	Phe	Ala 175	Leu
Trp	Asp	Ala	Leu 180	Asp	Leu	Gly	Asp	His 185	Val	Val	Leu	Val	Leu 190	His	Asp
Trp	Gly	Ser 195	Ala	Leu	Gly	Phe	Asp 200	Trp	Ala	Asn	Gln	His 205	Arg	Asp	Arg
Val	Gln 210	Gly	Ile	Ala	Phe	Met 215	Glu	Ala	Ile	Val	Thr 220	Pro	Met	Thr	Trp
Ala 225	Asp	Trp	Pro	Pro	Ala 230	Val	Arg	Gly	Val	Phe 235	Gln	Gly	Phe	Arg	Ser 240
Pro	Gln	Gly	Glu	Pro 245	Met	Ala	Leu	Glu	His 250	Asn	Ile	Phe	Val	Glu 255	Arg
Val	Leu	Pro	Gly 260	Ala	Ile	Leu	Arg	Gln 265	Leu	Ser	Asp	Glu	Glu 270	Met	Asn
His	Tyr	Arg 275	Arg	Pro	Phe	Val	Asn 280	Gly	Gly	Glu	Asp	Arg 285		Pro	Thr
Leu	Ser 290	Trp	Pro	Arg	Asn	Leu 295	Pro	Ile	Asp	Gly	Glu 300	Pro	Ala	Glu	Val
Val 305		Leu	Val	Asn			Arg			Leu 315		Glu	Thr	Asp	Met 320
Pro	Lys	Leu	Phe	11e 325	Asn	Ala	Glu	Pro	Gly 330		Ile	Ile	Thr	Gly 335	Arg
Ile	Arg	Asp	Tyr 340	Val	Arg	Ser	Trp	Pro 345		Gln	Thr	Glu	11e 350	Thr	Val
Pro	Gly	Val 355	His	Phe	Val	Gln	Glu 360	Asp	Ser	Asp	Gly	Val 365		. Ser	Trp
Ala	Gly 370	Ala	Arg	Gln	His	Arg 375	Arg	Pro	Gly	Ser	Ala 380		Ile	e Ser	Arg
Asp 385	Gln	Glu	Cys	Asp	Phe 390	Arg	Arg	Arg	Arg	Arg 395		Ala	Cys	Glr	1 Leu 400

Ile Arg Leu Pro Ala Pro Gly Arg Asp Ser Gln Gly Lys Gly His Gln 405 410 415

Ser Gln Pro Leu Pro Ser Gln Arg Gly Arg Gln Ile Tyr Val Ala Gly 420 425 430

Gln Arg Ser Ser Tyr Leu Pro Ser Glu Leu Val Ala Ala Phe Leu Trp 435 440 445

Ala Gln Phe Glu Glu Ala Glu Arg Ile Thr Arg Ile Arg Leu Asp Leu 450 455 460

Trp Asn Arg Tyr His Glu Ser Phe Glu Ser Leu Glu Gln Arg Gly Leu 465 470 475 480

Leu Arg Arg Pro Ile Ile Pro Gln Gly Cys Ser His Asn Ala Hi/s Met 485 490 495

Tyr Tyr Val Leu Leu Ala Pro Ser Ala Asp Arg Glu Glu Val Leu Ala . 500 505 510

Arg Leu Thr Ser Glu Gly Ile Gly Ala Val Phe His Tyr Val Pro Leu 515 520 525

His Asp Ser Pro Ala Gly Arg Arg 530 535

(2) INFORMATION FOR SEQ ID NO:157:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 284 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:157:

Asn Glu Ser Ala Pro Arg Ser Pro Met Leu Pro Ser Ala Arg Pro Arg

1 10 15

Tyr Asp Ala Ile Ala Val Leu Leu Asn Glu Met His Ala Gly His Cys 20 25 30

Asp Phe Gly Leu Val Gly Pro Ala Pro Asp Ile Val Thr Asp Ala Ala 35 40 45

Gly Asp Asp Arg Ala Gly Leu Gly Val Asp Glu Gln Phe Arg His Val 50 55 60

Gly Phe Leu Glu Pro Ala Pro Val Leu Val Asp Gln Arg Asp Asp Leu

65 70 75 80 Gly Gly Leu Thr Val Asp Trp Lys Val Ser Trp Pro Arg Gln Arg Gly 90 Ala Thr Val Leu Ala Ala Val His Glu Trp Pro Pro Ile Val Val His Phe Leu Val Ala Glu Leu Ser Gln Asp Arg Pro Gly Gln His Pro Phe 115 120 Asp Lys Asp Val Val Leu Gln Arg His Trp Leu Ala Leu Arg Arg Ser 135 Glu Thr Leu Glu His Thr Pro His Gly Arg Arg Pro Val Arg Pro Arg 145 155 His Arg Gly Asp Asp Arg Phe His Glu Arg Asp Pro Leu His Ser Val Ala Met Leu Val Ser Pro Val Glu Ala Glu Arg Arg Ala Pro Val Val 185 Gln His Gln Tyr His Val Val Ala Glu Val Glu Arg Ile Pro Glu Arg 195 Glu Gln Lys Val Ser Leu Leu Ala Ile Ala Ile Ala Val Gly Ser Arg 215 Trp Ala Glu Leu Val Arg Arg Ala His Pro Asp Gln Ile Ala Gly His 225 235 Gln Pro Ala Gln Pro Phe Gln Val Arg His Asp Val Ala Pro Gln Val 245 250 Arg Arg Arg Gly Val Ala Val Leu Lys Asp Asp Gly Val Thr Leu Ala 265 Phe Val Asp Ile Arg His Ala Leu Pro Gly Asp Phe 275 280

(2) INFORMATION FOR SEQ ID NO:158:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 264 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:158:

ATGAACATGT	CGTCGGTGGT	GGGTCGCAAG	GCCTTTGCGC	GATTCGCCGG	CTACTCCTCC	60
GCCATGCACG	CGATCGCCGG	TTTCTCCGAT	GCGTTGCGCC	AAGAGCTGCG	GGGTAGCGGA	120
ATCGCCGTCT	CGGTGATCCA	CCCGGCGCTG	ACCCAGACAC	CGCTGTTGGC	CAACGTCGAC	180
CCCGCCGACA	TGCCGCCGCC	GTTTCGCAGC	CTCACGCCCA	TTCCCGTTCA	CTGGGTCGCG	240
GCAGCGGTGC	TTGACGGTGT	GGCG				264

(2) INFORMATION FOR SEQ ID NO:159:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1171 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:159:

TAGTCGGCGA	CGATGACGTC	GCGGTCCAGG	CCGACCGCTT	CAAGCACCAG	CGCGACCACG	60
AAGCCGGTGC	GATCCTTACC	CGCGAAGCAG	TGGGTGAGCA	CCGGGCGTCC	GGCGGCAAGC	120
AGTGTGACGA	CACGATGTAG	CGCGCGCTGT	GCTCCATTGC	GCGTTGGGAA	TTGGCGATAC	180
TCGTCGGTCA	TGTAGCGGGT	GGCCGCGTCA	TTTATCGACT	GGCTGGATTC	GCCGGACTCG	240
CCGTTGGACC	CGTCATTGGT	TAGCAGCCTC	TTGAATGCGG	TTTCGTGCGG	CGCTGAGTCG	300
TCGGCGTCAT	CATCGGCGAG	GTCGGGGAAC	GGCAGCAGGT	GGACGTCGAT	GCCGTCCGGA	360
ACCCGTCCTG	GACCGCGGCG	GGCAACCTCC	CGGGACGACC	GCAGGTCGGC	AACGTCGGTG	420
ATCCCCAGCC	GGCGCAGCGT	TGCCCCTCGT	GCCGAATTCG	GCACGAGGCT	GGCGAGCCAC	480
CGGGCATCAC	CAAGCAACGC	TTGCCCAGTA	CGGATCGTCA	CTTCCGCATC	CGGCAGACCA	540
ATCTCCTCGC	CGCCCATCGT	CAGATCCCGC	TCGTGCGTTG	ACAAGAACGG	CCGCAGATGT	600
GCCAGCGGGT	ATCGGAGATT	GAACCGCGCA	CGCAGTTCTT	CAATCGCTGC	GCGCTGCCGC	660
ACTATTGGCA	CTTTCCGGCG	GTCGCGGTAT	TCAGCAAGCA	TGCGAGTCTC	GACGAACTCG	720
CCCCACGTAA	CCCACGGCGT	AGCTCCCGGC	GTGACGCGGA	GGATCGGCGG	GTGATCTTTG	780
CCGCCACGCT	CGTAGCCGTT	GATCCACCGC	TTCGCGGTGC	CGGCGGGGAG	GCCGATCAGC	840
TTATCGACCT	CGGCGTATGC	CGACGGCAAG	CTGGGCGCGT	TCGTCGAGGT	CAAGAACTCC	900
ACCATCGGCA	CCGGCACCAA	GGTGCCGCAC	CTGACCTACG	TCGGCGACGC	CGACATCGGC	960

GAGTACAGC	A ACATCGGCGC	CTCCAGCGTG	TTCGTCAACT	ACGACGGTAC	GTCCAAACGG	1020
CGCACCACC	G TCGGTTCGCA	CGTACGGACC	GGGTCCGACA	CCATGTTCGT	GGCCCCAGTA	1080
ACCATCGGC	G ACGGCGCGTA	TACCGGGGCC	GGCACAGTGG	TGCGGGAGGA	TGTCCCGCCG	1140
GGGGCGCTG	CAGTGTCGGC	GGGTCCGCAA	С			1171
(2) INFORM	MATION FOR S	EQ ID NO:160):			
(i) S	(B) TYPE: n	227 base pa ucleic acid DNESS: singl	airs			
				·	/	
(xi) S	EQUENCE DES	CRIPTION: SE	Q ID NO:160):		•
GCAAAGGCGG	CACCGGCGGG	GCCGGCATGA	ACAGCCTCGA	CCCGCTGCTA	GCCGCCCAAG	60
ACGGCGGCCA	AGGCGGCACC	GGCGGCACCG	GCGGCAACGC	CGGCGCCGGC	GGCACCAGCT	120
TCACCCAAGG	CGCCGACGGC	AACGCCGGCA	ACGGCGGTGA	CGGCGGGGTC	GGCGGCAACG	180
GCGGAAACGG	CGGAAACGGC	GCAGACAACA	CCACCACCGC	CGCCGCC		227
(2) INFORM	ATION FOR SE	EQ ID NO:161	:			
	EQUENCE CHAF (A) LENGTH: (B) TYPE: nu (C) STRANDED (D) TOPOLOGY	304 base pa cleic acid NESS: singl	irs			

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:161:

CCTCGCCACC	ATGGGCGGGC	AGGGCGGTAG	CGGTGGCGCC	GGCTCTACCC	CAGGCGCCAA	60
GGGCGCCCAC	GGCTTCACTC	CAACCAGCGG	CGGCGACGGC	GGCGACGGCG	GCAACGGCGG	120
CAACTCCCAA	GTGGTCGGCG	GCAACGGCGG	CGACGGCGGC	AATGGCGGCA	ACGGCGGCAG	180
CGCCGGCACG	GGCGGCAACG	GCGGCCGCGG	CGGCGACGGC	GCGTTTGGTG	GCATGAGTGC	240
CAACGCCACC	AACCCTGGTG	AAAACGGGCC	AAACGGTAAC	CCCGGCGGCA	ACGGTGGCGC	300

CGGC 304

(2) INFORMATION FOR SEQ ID NO:162:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1439 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:162:

GTGGGACGCT	GCCGAGGCTG	TATAACAAGG	ACAACATCGA	CCAGCGCCGG	CTCGGTGAGC	60
TGATCGACCT	ATTTAACAGT	GCGCGCTTCA	GCCGGCAGGG	CGAGCACCGC	GCCCGGGATC	120
TGATGGGTGA	GGTCTACGAA	TACTTCCTCG	GCAATTTCGC	TCGCGCGGAA	GGGAAGCGGG	180
GTGGCGAGTT	CTTTACCCCG	CCCAGCGTGG	TCAAGGTGAT	CGTGGAGGTG	CTGGAGCCGT	240
CGAGTGGGCG	GGTGTATGAC	CCGTGCTGCG	GTTCCGGAGG	CATGTTTGTG	CAGACCGAGA	300
AGTTCATCTA	CGAACACGAC	GGCGATCCGA	AGGATGTCTC	GATCTATGGC	CAGGAAAGCA	360
TTGAGGAGAC	CTGGCGGATG	GCGAAGATGA	ACCTCGCCAT	CCACGGCATC	GACAACAAGG	420
GGCTCGGCGC	CCGATGGAGT	GATACCTTCG	CCCGCGACCA	GCACCCGGAC	GTGCAGATGG	480
ACTACGTGAT	GGCCAATCCG	CCGTTCAACA	TCAAAGACTG	GGCCCGCAAC	GAGGAAGACC	540
CACGCTGGCG	CTTCGGTGTT	CCGCCCGCCA	ATAACGCCAA	CTACGCATGG	ATTCAGCACA	600
TCCTGTACAA	CTTGGCGCCG	GGAGGTCGGG	CGGGCGTGGT	GATGGCCAAC	GGGTCGATGT	660
CGTCGAACTC	CAACGGCAAG	GGGGATATTC	GCGCGCAAAT	CGTGGAGGCG	GATTTGGTTT	720
CCTGCATGGT	CGCGTTACCC	ACCCAGCTGT	TCCGCAGCAC	CGGAATCCCG	GTGTGCCTGT	780
GGTTTTTCGC	CAAAAACAAG	GCGGCAGGTA	AGCAAGGGTC	TATCAACCGG	TGCGGGCAGG	840
TGCTGTTCAT	CGACGCTCGT	GAACTGGGCG	ACCTAGTGGA	CCGGGCCGAG	CGGGCGCTGA	900
CCAACGAGGA	GATCGTCCGC	ATCGGGGATA	CCTTCCACGC	GAGCACGACC	ACCGGCAACG	960
CCGGCTCCGG	TGGTGCCGGC	GGTAATGGGG	GCACTGGCCT	CAACGGCGCG	GGCGGTGCTG	1020
GCGGGGCCGG	CGGCAACGCG	GGTGTCGCCG	GCGTGTCCTT	CGGCAACGCT	GTGGGCGGCG	1080
ACGGCGGCAA	CGGCGGCAAC	GGCGGCCACG	GCGGCGACGG	CACGACGGGC	GGCGCCGGCG	1140
GCAAGGGCGG	CAACGGCAGC	AGCGGTGCCG	CCAGCGGCTC	AGGCGTCGTC	AACGTCACCG	1200

CCGGCCACGG	CGGCAACGGC	GGCAATGGCG	GCAACCCCC	Chhacaamaa	GCGGGCGCCG	
						1260
GCGGCCAGGG	CGGTGCCGGC	GGCAGCGCCG	GCAACGGCGG	CCACGGCGGC	GGTGCCACCG	1320
GCGGCGCCAG	CGGCAAGGGC	GGCAACGGCA	CCAGCGGTGC	CGCCAGCGGC	TCAGGCGTCA	1380
						1380
TCAACGTCAC	CGCCGGCCAC	GGCGGCAACG	GCGGCAATGG	CCGCAACGGC	GGCAACGGC	1439
(2) INFORMA	TION FOR SE	O ID NO-163	•			

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 329 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:163:

(2) INFORMATION FOR SEQ ID NO:164:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 80 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:164:

GCAACGGTGG CAACGGCGGC ACCAGCACGA CCGTGGGGAT GGCCGGAGGT AACTGTGGTG 60
CCGCCGGGCT GATCGGCAAC 80

(2) INFORMATION FOR SEQ ID NO:165:

PCT/US97/18214

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 392 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:165:

GGGCTGTGTC	GCACTCACAC	CGCCGCATTC	GGCGACGTTG	GCCGCCCAAT	ATCCAGCTCA	60
AGGCCTACTA	CTTACCGTCG	GAGGACCGCC	GCATCAAGGT	GCGGGTCAGC	GCCCAAGGAA	120
TCAAGGTCAT	CGACCGCGAC	GGGCATCGAG	GCCGTCGTCG	CGCGGCTCGG	GCAGGATCCG	180
CCCCGGCGCA	CTTCGCGCGC	CAAGCGGGCT	CATCGCTCCG	AACGGCGGCG	ATCCTGTGAG	240
CACAACTGAT	GGCGCGCAAC	GAGATTCGTC	CAATTGTCAA	GCCGTGTTCG	ACCGCAGGGA	300
CCGGTTATAC	GTATGTCAAC	CTATGTCACT	CGCAAGAACC	GGCATAACGA	TCCCGTGATC	360
CGCCGACAGC	CCACGAGTGC	AAGACCGTTA	CA			392

(2) INFORMATION FOR SEQ ID NO:166:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 535 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:166:

ACCGGCGCCA	CCGGCGGCAC	CGGGTTCGCC	GGTGGCGCCG	GCGGGGCCGG	CGGGCAGGGC	60
GGTATCAGCG	GTGCCGGCGG	CACCAACGGC	TCTGGTGGCG	CTGGCGGCAC	CGGCGGACAA	120
GGCGGCGCCG	GGGGCGCTGG	CGGGGCCGGC	GCCGATAACC	CCACCGGCAT	CGGCGGCGCC	180
GGCGGCACCG	GCGGCACCGG	CGGAGCGGCC	GGAGCCGGCG	GGGCCGGTGG	CGCCATCGGT	240
ACCGGCGGCA	CCGGCGGCGC	GGTGGGCAGC	GTCGGTAACG	CCGGGATCGG	CGGTACCGGC	300
GGTACGGGTG	GTGTCGGTGG	TGCTGGTGGT	GCAGGTGCGG	CTGCGGCCGC	TGGCAGCAGC	360
GCTACCGGTG	GCGCCGGGTT	CGCCGGCGGC	GCCGGCGGAG	AAGGCGGACC	GGGCGGCAAC	420
AGCGGTGTGG	GCGGCACCAA	CGGCTCCGGC	GGCGCCGGCG	GTGCAGGCGG	CAAGGCCGC	480

ACCGGAGGTG	CCGGCGGGTC	CGGCGCGGAC	AACCCCACCG	GTGCTGGTTT	CGCCG	535

(2) INFORMATION FOR SEQ ID NO:167:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 690 base pairs

 - (B) TYPE: nucleic acid(C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:167:

CCGACGTCGC	CGGGGCGATA	CGGGGGTCAC	CGACTACTAC	ATCATCCGCA	CCGAGAATCG	60
GCCGCTGCTG	CAACCGCTGC	GGGCGGTGCC	GGTCATCGGA	GATCCGCTGG	CCGACCTGAT	120
CCAGCCGAAC	CTGAAGGTGA	TCGTCAACCT	GGGCTACGGC	GACCCGAACT	ACGGCTACTC	180
GACGAGCTAC	GCCGATGTGC	GAACGCCGTT	CGGGCTGTGG	CCGAACGTGC	CGCCTCAGGT	240
CATCGCCGAT	GCCCTGGCCG	CCGGAACACA	AGAAGGCATC	CTTGACTTCA	CGGCCGACCT	300
GCAGGCGCTG	TCCGCGCAAC	CGCTCACGCT	CCCGCAGATC	CAGCTGCCGC	AACCCGCCGA	360
TCTGGTGGCC	GCGGTGGCCG	CCGCACCGAC	GCCGGCCGAG	GTGGTGAACA	CGCTCGCCAG	420
GATCATCTCA	ACCAACTACG	ÇCGTCCTGCT	GCCCACCGTG	GACATCGCCC	TCGCCTGGTC	480
ACCACCCTGC	CGCTGTACAC	CACCCAACTG	TTCGTCAGGC	AACTCGCTGC	GGGCAATCTG	540
ATCAACGCGA	TCGGCTATCC	CCTGGCGGCC	ACCGTAGGTT	TAGGCACGAT	CGATAGCGGG	600
CGGCGTGGAA	TTGCTCACCC	TCCTCGCGGC	GGCCTCGGAC	ACCGTTCGAA	ACATCGAGGG	660
CCTCGTCACC	TAACGGATTC	CCGACGGCAT				690

(2) INFORMATION FOR SEQ ID NO:168:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 407 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

ACGGTGACGG	CGGTACTGGC	GGCGGCCACG	GCGGCAACGG	CGGGAATCCC	GGGTGGCTCT	60
TGGGCACAGC	CGGGGGTGGC	GGCAACGGTG	GCGCCGGCAG	CACCGGTACT	GCAGGTGGCG	120
GCTCTGGGGG	CACCGGCGGC	GACGGCGGGA	CCGGCGGGCG	TGGCGGCCTG	TTAATGGGCG	180
CCGGCGCCGG	CGGGCACGGT	GGCACTGGCG	GCGCGGGCGG	TGCCGGTGTC	GACGGTGGCG	240
GCGCCGGCGG	GGCCGGCGGG	GCCGGCGCA	ACGGCGGCGC	CGGGGGTCAA	GCCGCCCTGC	300
TGTTCGGGCG	CGGCGGCACC	GGCGGAGCCG	GCGGCTACGG	CGGCGATGGC	GGTGGCGGCG	360
GTGACGGCTT	CGACGGCACG	ATGGCCGGCC	TGGGTGGTAC	CGGTGGC		407

(2) INFORMATION FOR SEQ ID NO:169:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 468 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:169:

GATCGGTCAG	CGCATCGCCC	TCGGCGGCAA	GCGATTCCGC	GGTCTCACCG	AAGAACATCG	60
TGCACGCGGC	GGCGCGGACC	AGCCCGCTGC	GCTGCGGCGC	GTCGAACGCC	TCCAGCAGGC	120
ACAGCCAGTC	CTTGGCGGCC	TGCGAGGCGA	ACACGTCGGT	GTCACCGGTG	TAGATCGCCG	180
GGATGCCCGC	CTCCGCCAAC	GCATTCCGGC	ACGCCCGCGC	GTCTTTGTGA	TGCTCGACGA	240
TCACCGCGAT	GTCTGCGGCC	ACCACGGGCC	GCCCGGCGAA	GGTGGCCCCG	CTGGCCAGTA	300
GCGCCGCGAC	GTCGGCGGCC	AGGTCGTCGG	GGATGTGCCG	GCGCAGCGCT	CCGGCGCGAC	360
GCCCGAAAAA	CGACCCCTCA	CCCAGCTGGG	TCCCGCTGGC	ATATCCCTTG	CCGTCCTGGG	420
CGATATTGGA	CGCGCATGCC	CCGACCGCGT	ACAGGCCGGC	CACCACCG		468

(2) INFORMATION FOR SEQ ID NO:170:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 219 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE	DESCRIPTION:	SEQ	ID	NO:170:
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GGTGGTAACG GCGGCCAGGG TGGCATCGG	c gececceece	AGAGAGGCGC	CGACGGCGCC	60
GGCCCCAATG CTAACGGCGC AAACGGCGA	G AACGGCGGTA	GCGGTGGTAA	CGGTGGCGAC	120
GGCGGCGCG GCGCAATGG CGGCGCGGG	C GGCAACGCGC	AGGCGGCCGG	GTACACCGAC	180
GGCGCCACGG GCACCGGCGG CGACGGCGG	C AACGGCGGC	•		219

(2) INFORMATION FOR SEQ ID NO:171:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 494 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:171:

TAGCTCCGGC	GAGGGCGGCA	AGGGCGGCGA	CGGTGGCCAC	GGCGGTGACG	GCGTCGGCGG	60
CAACAGTTCC	GTCACCCAAG	GCGGCAGCGG	CGGTGGCGGC	GGCGCCGGCG	GCGCCGGCGG	120
CAGCGGCTTT	TTCGGCGGCA	AGGGCGGCTT	CGGCGGCGAC	GGCGGTCAGG	GCGGCCCCAA	180
CGGCGGCGGT	ACCGTCGGCA	CCGTGGCCGG	TGGCGGCGGC	AACGGCGGTG	TCGGCGGCCG	240
GGGCGGCGAC	GGCGTCTTTG	CCGGTGCCGG	CGGCCAGGGC	GGCCTCGGTG	GGCAGGGCGG	300
CAATGGCGGC	GGCTCCACCG	GCGGCAACGG	CGGCCTTGGC	GGCGCGGGCG	GTGGCGGAGG	360
CAACGCCCCG	GCTCGTGCCG	AATCCGGGCT	GACCATGGAC	AGCGCGGCCA	AGTTCGCTGC	420
CATCGCATCA	GGCGCGTACT	GCCCCGAACA	CCTGGAACAT	CACCCGAGTT	AGCGGGGCGC	480
ATTTCCTGAT	CACC					494

(2) INFORMATION FOR SEQ ID NO:172:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 220 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

PCT/US97/18214

185

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:172:	
GGGCCGGTGG TGCCGCGGGC CAGCTCTTCA GCGCCGGAGG CGCGGCGGGT GCCGTTGGGG	60
TTGGCGGCAC CGGCGGCCAG GGTGGGGCTG GCGGTGCCGG AGCGGCCGGC GCCGACGCCC	120
CCGCCAGCAC AGGTCTAACC GGTGGTACCG GGTTCGCTGG CGGGGCCGGC GGCGTCGGCG	180
GCCAGAGCGG CAACGCCATT GCCGGCGGCA TCAACGGCTC	220
(2) INFORMATION FOR SEQ ID NO:173:	·

(i) SEQUENCE CHARACTERISTICS:

- - (A) LENGTH: 388 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:173:

ATGGCGGCAA CGGGGGCCCC GGCGGTGCTG GCGGGGCCGG CGACTACAAT TTCCAACGGC 60 GGGCAGGGTG GTGCCGGCGG CCAAGGCGGC CAAGGCGGCC TGGGCGGGGC AAGCACCACC 120 TGATCGGCCT AGCCGCACCC GGGAAAGCCG ATCCAACAGG CGACGATGCC GCCTTCCTTG 180 CCGCGTTGGA CCAGGCCGGC ATCACCTACG CTGACCCAGG CCACGCCATA ACGGCCGCCA 240 AGGCGATGTG TGGGCTGTGT GCTAACGGCG TAACAGGTCT ACAGCTGGTC GCGGACCTGC 300 GGGACTACAA TCCCGGGCTG ACCATGGACA GCGCGGCCAA GTTCGCTGCC ATCGCATCAG 360 388 GCGCGTACTG CCCCGAACAC CTGGAACA

(2) INFORMATION FOR SEQ ID NO:174:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 400 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:174:

GCAAAGGCGG CACCGGCGGG GCCGGCATGA ACAGCCTCGA CCCGCTGCTA GCCGCCCAAG 60 ACGGCGGCCA AGGCGGCACC GGCGCCACCG GCGCCAACGC CGGCGCCGGC GGCACCAGCT 120

TCACCCAAGG	CGCCGACGGC	AACGCCGGCA	ACGGCGGTGA	CGGCGGGGTC	GGCGGCAACG	180
GCGGAAACGG	CGGAAACGGC	GCAGACAACA	CCACCACCGC	CGCCGCCGGC	ACCACAGGCG	240
GCGACGGCGG	GGCCGGCGGG	GCCGGCGGAA	CCGGCGGAAC	CGGCGGAGCC	GCCGGCACCG	300
GCACCGGCGG	CCAACAAGGC	AACGGCGGCA	ACGGCGGCAC	CGGCGGCAAA	GGCGGCACCG	360
GCGGCGACGG	TGCACTCTCA	GGCAGCACCG	GTGGTGCCGG			400

(2) INFORMATION FOR SEQ ID NO:175:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 538 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:175:

GGCAACGGCG	GCAACGGCGG	CATCGCCGGC	ATTGGGCGGC	AACGGCGTTC	CGGGACGGGC	60
AGCGGCAACG	GCGGCCAACG	GCGGCAGCGG	CGGCAACGGC	GGCAACGCCG	GCATGGGCGG	120
CAACAGCGGC	ACCGGCAGCG	GCGACGGCGG	TGCCGGCGGG	AACGGCGGCG	CGGCGGCAC	180
GGGCGGCACC	GGCGGCGACG	GCGGCCTCAC	CGGTACTGGC	GGCACCGGCG	GCAGCGGTGG	240
CACCGGCGGT	GACGGCGGTA	ACGGCGGCAA	CGGAGCAGAT	AACACCGCAA	ACATGACTGC	300
GCAGGCGGGC	GGTGACGGTG	GCAACGGCGG	CGACGGTGGC	TTCGGCGGCG	GGGCCGGGGC	360
CGGCGGCGGT	GGCTTGACCG	CTGGCGCCAA	CGGCACCGGC	GGGCAAGGCG	GCGCCGGCGG	420
CGATGGCGGC	AACGGGGCCA	TCGGCGGCCA	CGGCCCACTC	ACTGACGACC	CCGGCGGCAA	480
CGGGGGCACC	GGCGGCAACG	GCGGCACCGG	CGGCACCGGC	GGCGCGGGCA	TCGGCAGC	538

(2) INFORMATION FOR SEQ ID NO:176:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 239 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:176:

GGGCCGGTGG	TGCCGCGGGC	CAGCTCTTCA	GCGCCGGAGG	CGCGGCGGGT	GCCGTTGGGG	60
TTGGCGGCAC	CGGCGGCCAG	GGTGGGGCTG	GCGGTGCCGG	AGCGGCCGGC	GCCGACGCCC	120
CCGCCAGCAC	AGGTCTAACC	GGTGGTACCG	GGTTCGCTGG	CGGGGCCGGC	GGCGTCGGCG	180
GCCACGGCGG	CAACGCCATT	GCCGGCGGCA	TCAACGGCTC	CGGTGGTGCC	GGCGGCACC	239

(2) INFORMATION FOR SEQ ID NO:177:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 985 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:177:

AGCAGCGCTA	CCGGTGGCGC	CGGGTTCGCC	GGCGGCGCCG	GCGGAGAAGG	CGGAGCGGC	60
GGCAACAGCG	GTGTGGGCGG	CACCAACGGC	TCCGGCGGCG	CCGGCGGTGC	AGGCGGCAAG	120
GGCGGCACCG	GAGGTGCCGG	CGGGTCCGGC	GCGGACAACC	CCACCGGTGC	TGGTTTCGCC	180
GGTGGCGCCG	GCGGCACAGG	TGGCGCGGCC	GGCGCCGGCG	GGGCCGGCGG	GGCGACCGGT	240
ACCGGCGGCA	CCGGCGGCGT	TGTCGGCGCC	ACCGGTAGTG	CAGGCATCGG	CGGGGCCGGC	300
GGCCGCGGCG	GTGACGGCGG	CGATGGGGCC	AGCGGTCTCG	GCCTGGGCCT	CTCCGGCTTT	360
GACGGCGGCC	AAGGCGGCCA	AGGCGGGGCC	GGCGGCAGCG	CCGGCGCCGG	CGGCATCAAC	420
GGGGCCGGCG	GGGCCGGCGG	CAACGGCGGC	GACGGCGGGG	ACGGCGCAAC	CGGTGCCGCA	480
GGTCTCGGCG	ACAACGGCGG	GGTCGGCGGT	GACGGTGGGG	CCGGTGGCGC	CGCCGGCAAC	540
GGCGGCAACG	CGGGCGTCGG	CCTGACAGCC	AAGGCCGGCG	ACGGCGGCGC	CGCGGGCAAT	600
GGCGGCAACG	GGGGCGCCGG	CGGTGCTGGC	GGGGCCGGCG	ACAACAATTT	CAACGGCGGC	660
CAGGGTGGTG	CCGGCGGCCA	AGGCGGCCAA	GGCGGCTTGG	GCGGGGCAAG	CACCACCTGA	720
TCGGCCTAGC	CGCACCCGGG	AAAGCCGATC	CAACAGGCGA	CGATGCCGCC	TTCCTTGCCG	780
CGTTGGACCA	GGCCGGCATC	ACCTACGCTG	ACCCAGGCCA	CGCCATAACG	GCCGCCAAGG	840
CGATGTGTGG	GCTGTGTGCT	AACGGCGTAA	CAGGTCTACA	GCTGGTCGCG	GACCTGCGGG	900
AATACAATCC	CGGGCTGACC	ATGGACAGCG	CGGCCAAGTT	CGCTGCCATC	GCATCAGGCG	960

CGTACTGCCC CGAACACCTG GAACA

985

(2) INFORMATION FOR SEQ ID NO:178:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2138 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:178:

CGGCACGAGG ATCGGTACCC CGCGGCATCG GCAGCTGCCG ATTCGCCGGG TTTCCCCACC	60
CGAGGAAAGC CGCTACCAGA TGGCGCTGCC GAAGTAGGGC GATCCGTTCG CGATGCCGGC	120
ATGAACGGGC GGCATCAAAT TAGTGCAGGA ACCTTTCAGT TTAGCGACGA TAATGGCTAT	180
AGCACTAAGG AGGATGATCC GATATGACGC AGTCGCAGAC CGTGACGGTG GATCAGCAAG	240
AGATTTTGAA CAGGGCCAAC GAGGTGGAGG CCCCGATGGC GGACCCACCG ACTGATGTCC	300
CCATCACACC GTGCGAACTC ACGGCGGCTA AAAACGCCGC CCAACAGCTG GTATTGTCCG	. 360
CCGACAACAT GCGGGAATAC CTGGCGGCCG GTGCCAAAGA GCGGCAGCGT CTGGCGACCT	420
CGCTGCGCAA CGCGGCCAAG GCGTATGGCG AGGTTGATGA GGAGGCTGCG ACCGCGCTGG	480
ACAACGACGG CGAAGGAACT GTGCAGGCAG AATCGGCCGG GGCCGTCGGA GGGGACAGTT	540
CGGCCGAACT AACCGATACG CCGAGGGTGG CCACGGCCGG TGAACCCAAC TTCATGGATC	600
TCAAAGAAGC GGCAAGGAAG CTCGAAACGG GCGACCAAGG CGCATCGCTC GCGCACTTTG	660
CGGATGGGTG GAACACTTTC AACCTGACGC TGCAAGGCGA CGTCAAGCGG TTCCGGGGGT	720
TTGACAACTG GGAAGGCGAT GCGGCTACCG CTTGCGAGGC TTCGCTCGAT CAACAACGGC	780
AATGGATACT CCACATGGCC AAATTGAGCG CTGCGATGGC CAAGCAGGCT CAATATGTCG	840
CGCAGCTGCA CGTGTGGGCT AGGCGGGAAC ATCCGACTTA TGAAGACATA GTCGGGCTCG	900
AACGGCTTTA CGCGGAAAAC CCTTCGGCCC GCGACCAAAT TCTCCCGGTG TACGCGGAGT	960
ATCAGCAGAG GTCGGAGAAG GTGCTGACCG AATACAACAA CAAGGCAGCC CTGGAACCGG	1020
TAAACCCGCC GAAGCCTCCC CCCGCCATCA AGATCGACCC GCCCCCGCCT CCGCAAGAGC	1080
AGGGATTGAT CCCTGGCTTC CTGATGCCGC CGTCTGACGG CTCCGGTGTG ACTCCCGGTA	1140

CCGGGATGCC	AGCCGCACCG	ATGGTTCCGC	CTACCGGATC	GCCGGGTGGT	GGCCTCCCGG	1200
CTGACACGGC	GGCGCAGCTG	ACGTCGGCTG	GGCGGGAAGC	CGCAGCGCTG	TCGGGCGACG	1260
TGGCGGTCAA	AGCGGCATCG	CTCGGTGGCG	GTGGAGGCGG	CGGGGTGCCG	TCGGCGCCGT	1320
TGGGATCCGC	GATCGGGGGC	GCCGAATCGG	TGCGGCCCGC	TGGCGCTGGT	GACATTGCCG	1380
GCTTAGGCCA	GGGAAGGCC	GGCGGCGGCG	CCGCGCTGGG	CGGCGGTGGC	ATGGGAATGC	1440
CGATGGGTGC	CGCGCATCAG	GGACAAGGGG	GCGCCAAGTC	CAAGGGTTCT	CAGCAGGAAG	1500
ACGAGGCGCT	CTACACCGAG	GATCGGGCAT	GGACCGAGGC	CGTCATTGGT	AACCGTCGGC	1560
GCCAGGACAG	TAAGGAGTCG	AAGTGAGCAT	GGACGAATTG	GACCCGCATG	TCGCCCGGGC	1620
GTTGACGCTG	GCGGCGCGGT	TTCAGTCGGC	CCTAGACGGG	ACGCTCAATC	AGATGAACAA	1680
CGGATCCTTC	CGCGCCACCG	ACGAAGCCGA	GACCGTCGAA	GTGACGATCA	ATGGGCACCA	1740
GTGGCTCACC	GGCCTGCGCA	TCGAAGATGG	TTTGCTGAAG	AAGCTGGGTG	CCGAGGCGGT	1800
GGCTCAGCGG	GTCAACGAGG	CGCTGCACAA	TGCGCAGGCC	GCGGCGTCCG	CGTATAACGA	1860
CGCGGCGGGC	GAGCAGCTGA	CCGCTGCGTT	ATCGGCCATG	TCCCGCGCGA	TGAACGAAGG	1920
AATGGCCTAA	GCCCATTGTT	GCGGTGGTAG	CGACTACGCA	CCGAATGAGC	GCCGCAATGC	1980
GGTCATTCAG	CGCGCCCGAC	ACGGCGTGAG	TACGCATTGT	CAATGTTTTG	ACATGGATCG	2040
GCCGGGTTCG	GAGGGCGCCA	TAGTCCTGGT	CGCCAATATT	GCCGCAGCTA	GCTGGTCTTA	2100
GGTTCGGTTA	CGCTGGTTAA	TTATGACGTC	CGTTACCA			2138

(2) INFORMATION FOR SEQ ID NO:179:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 460 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:179:

Met Thr Gln Ser Gln Thr Val Thr Val Asp Gln Gln Glu Ile Leu Asn 1 5 10 15

Arg Ala Asn Glu Val Glu Ala Pro Met Ala Asp Pro Pro Thr Asp Val 20 25 30

Pro Ile Thr Pro Cys Glu Leu Thr Ala Ala Lys Asn Ala Ala Gln Gln

			35						4 ()						45			
L€	eu V 5	al O	Le	u Se	er A	la	Asp	As 55	n Me	et	Arc	g Gl	u	Tyr	Lei 60	ı Al	a Al	a Gl	y Ala
Ly 65	rs G	lu	Ar	g Gl	ln A	rg	Leu 70	Al	a Th	ır	Sea	r Le		Arg 75	Ası	Al.	a Al	a Ly	s Ala 80
Ту	r G	ly	Glı	u Va	al A 8.	sp 5	Glu	Gl	u Al	a	Ala	Th 90	r.	Ala	Leu	ı Ası	o Ası	n Ası 95	o Gly
Gl	u Gi	lу	Thr	• Va 10	1 G	ln i	Ala	Glı	ມ Se	r	Ala 105	G1	у .	Ala	Val	Gly	y Gly		Ser
Se.	r Al	a	Glu 115	Le	u Ti	ır i	Asp	Th:	Pr 12	0	Arg	Va.	1 1	Ala	Thr	Ala 125		/ Glu	Pro
Ası	n Ph 13	ie :	Met	. As	p Le	eu I	Lys	Glu 135	a Al	a	Ala	Ar	g 1	Lys	Leu 140	Glu	Thr	Gly	Asp
Glr 145	n Gl	У	Ala	Se.	r Le	u A	11a .50	His	Phe	е.	Ala	Ası		31 y 155	Trp	Asn	Thr	Phe	2 Asn 160
Leu	Th	r]	Leu	Glı	n Gl 16	у А 5	sp	Val	Lys	S ,	Arg	Phe 170		Arg	Gly	Phe	Asp	Asn 175	Trp
				TR	J					-	185						190		Arg
		1	95						200)						205			Gln
	210	,						215							220				Pro
225						2.	30						2	35			Glu		240
					245)						250				ė	Gln	255	_
Ser	Glu	ı L	ys	Val 260	Leu	Tł	ır (Glu	Tyr	A 2	sn 65	Asn	L	ys .	Ala	Ala	Leu 270	Glu	Pro
Val	Asn	P: 2"	ro 75	Pro	Lys	Pr	o E	?ro	Pro 280	A	la	Ile	Ly	ys :	Ile	Asp 285	Pro	Pro	Pro
Pro	Pro 290	G]	ln (Glu	Gln	Gl	у I 2	eu 195	Ile	P	ro	Gly	Pł		Leu 300	Met	Pro	Pro	Ser
Asp 305	Gly	Se	er (Gly	Val	Th 31	r P O	ro	Gly	T	hr (Gly	M∈ 31		Pro	Ala	Ala	Pro	Met 320
Val	Pro	Pr	r o	hr	Gly 325	Se	r P	ro	Gly	G.		Gly 330	Le	eu E	ro .	Ala	Asp	Thr 335	Ala

Ala Gln Leu Thr Ser Ala Gly Arg Glu Ala Ala Ala Leu Ser Gly Asp 340 345 350

Val Ala Val Lys Ala Ala Ser Leu Gly Gly Gly Gly Gly Gly Val 355 360 365

Pro Ser Ala Pro Leu Gly Ser Ala Ile Gly Gly Ala Glu Ser Val Arg 370 375 380

Pro Ala Gly Ala Gly Asp Ile Ala Gly Leu Gly Gln Gly Arg Ala Gly 385 390 395 400

Gly Gly Ala Ala Leu Gly Gly Gly Gly Met Gly Met Pro Met Gly Ala 405 410 415

Ala His Gln Gly Gln Gly Gly Ala Lys Ser Lys Gly Ser Gln Gln Glu 420 425 . 430

Asp Glu Ala Leu Tyr Thr Glu Asp Arg Ala Trp Thr Glu Ala Val Ile 435 440 445

Gly Asn Arg Arg Gln Asp Ser Lys Glu Ser Lys 450 455 460

(2) INFORMATION FOR SEQ ID NO:180:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 277 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:180:

Ala Gly Asn Val Thr Ser Ala Ser Gly Pro His Arg Phe Gly Ala Pro
1 10 15

Asp Arg Gly Ser Gln Arg Arg Arg His Pro Ala Ala Ser Thr Ala 20 25 30

Thr Glu Arg Cys Arg Phe Asp Arg His Val Ala Arg Gln Arg Cys Gly 35 40 45

Phe Pro Pro Ser Arg Arg Gln Leu Arg Arg Arg Val Ser Arg Glu Ala

Thr Thr Arg Arg Ser Gly Arg Arg Asn His Arg Cys Gly Trp His Pro 65 70 75 80

Gly Thr Gly Ser His Thr Gly Ala Val Arg Arg Arg His Gln Glu Ala

ì

192

90 85 95 Arg Asp Gln Ser Leu Leu Leu Arg Arg Gly Arg Val Asp Leu Asp 100 105 Gly Gly Gly Arg Leu Arg Arg Val Tyr Arg Phe Gln Gly Cys Leu Val 120 Val Val Phe Gly Gln His Leu Leu Arg Pro Leu Leu Ile Leu Arg Val 135 130 His Arq Glu Asn Leu Val Ala Gly Arg Arg Val Phe Arg Val Lys Pro 150 155 Phe Glu Pro Asp Tyr Val Phe Ile Ser Arg Met Phe Pro Pro Ser Pro His Val Gln Leu Arg Asp Ile Leu Ser Leu Leu Gly His Arg Ser Ala 185 Gln Phe Gly His Val Glu Tyr Pro Leu Pro Leu Leu Ile Glu Arg Ser 200 Leu Ala Ser Gly Ser Arg Ile Ala Phe Pro Val Val Lys Pro Pro Glu 215 Pro Leu Asp Val Ala Leu Gln Arg Gln Val Glu Ser Val Pro Pro Ile 235 230 Arg Lys Val Arg Glu Arg Cys Ala Leu Val Ala Arg Phe Glu Leu Pro 250 Cys Arg Phe Phe Glu Ile His Glu Val Gly Phe Thr Gly Arg Gly His 265 Pro Arg Arg Ile Gly 275

(2) INFORMATION FOR SEQ ID NO:181:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 192 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:181:

Arg Val Ala Ala Ser Phe Ile Asp Trp Leu Asp Ser Pro Asp Ser Pro 1 5 10 15

- Leu Asp Pro Ser Leu Val Ser Ser Leu Leu Asn Ala Val Ser Cys Gly 25 Ala Glu Ser Ser Ala Ser Ser Ser Ala Arg Ser Gly Asn Gly Ser Arg 40 Trp Thr Ser Met Pro Ser Gly Thr Arg Pro Gly Pro Arg Arg Ala Thr 55 50 Ser Arg Asp Asp Arg Arg Ser Ala Thr Ser Val Ile Pro Ser Arg Arg 70 Ser Val Ala Pro Arg Ala Glu Phe Gly Thr Arg Leu Ala Ser His Arg 90 Ala Ser Pro Ser Asn Ala Cys Pro Val Arg Ile Val Thr Ser Ala Ser 105 Gly Arg Pro Ile Ser Ser Pro Pro Ile Val Arg Ser Arg Ser Cys Val 120 Asp Lys Asn Gly Arg Arg Cys Ala Ser Gly Tyr Arg Arg Leu Asn Arg 130 Ala Arg Ser Ser Ser Ile Ala Ala Arg Cys Arg Thr Ile Gly Thr Phe 155 150 Arg Arg Ser Arg Tyr Ser Ala Ser Met Arg Val Ser Thr Asn Ser Pro 170 165 His Val Thr His Gly Val Ala Pro Gly Val Thr Arg Arg Ile Gly Gly 185 180
- (2) INFORMATION FOR SEQ ID NO:182:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 196 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:182:
 - Gln Glu Arg Pro Gln Met Cys Gln Arg Val Ser Glu Ile Glu Pro Arg
 - Thr Gln Phe Phe Asn Arg Cys Ala Leu Pro His Tyr Trp His Phe Pro 20 25 30
 - Ala Val Ala Val Phe Ser Lys His Ala Ser Leu Asp Glu Leu Ala Pro

45 40 35 Arg Asn Pro Arg Arg Ser Ser Arg Arg Asp Ala Glu Asp Arg Arg Val Ile Phe Ala Ala Thr Leu Val Ala Val Asp Pro Pro Leu Arg Gly Ala Gly Glu Ala Asp Gln Leu Ile Asp Leu Gly Val Cys Arg Arg Gln Ala Gly Arg Val Arg Arg Gly Gln Glu Leu His His Arg His Arg His Gln Gly Ala Ala Pro Asp Leu Arg Arg Arg Arg Arg His Arg Arg Val 120 Gln Gln His Arg Arg Leu Gln Arg Val Arg Gln Leu Arg Arg Tyr/ Val 135 Gln Thr Ala His His Arg Arg Phe Ala Arg Thr Asp Arg Val Arg His 150 His Val Arg Gly Pro Ser Asn His Arg Arg Arg Val Tyr Arg Gly 175 165 Arg His Ser Gly Ala Gly Gly Cys Pro Ala Gly Gly Ala Gly Ser Val Gly Gly Ser Ala

(2) INFORMATION FOR SEQ ID NO:183:

195

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 311 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:183:

Val Arg Cys Gly Thr Leu Val Pro Val Pro Met Val Glu Phe Leu Thr 1 5 10 15

Ser Thr Asn Ala Pro Ser Leu Pro Ser Ala Tyr Ala Glu Val Asp Lys 20 25 30

Leu Ile Gly Leu Pro Ala Gly Thr Ala Lys Arg Trp Ile Asn Gly Tyr 35 40 45

- Glu Arg Gly Gly Lys Asp His Pro Pro Ile Leu Arg Val Thr Pro Gly 55 Ala Thr Pro Trp Val Thr Trp Gly Glu Phe Val Glu Thr Arg Met Leu Ala Glu Tyr Arg Asp Arg Arg Lys Val Pro Ile Val Arg Gln Arg Ala 90 85 Ala Ile Glu Glu Leu Arg Ala Arg Phe Asn Leu Arg Tyr Pro Leu Ala 105 100 His Leu Arg Pro Phe Leu Ser Thr His Glu Arg Asp Leu Thr Met Gly 120 Gly Glu Glu Ile Gly Leu Pro Asp Ala Glu Val Thr Ile Arg Thr Gly 140 135 130 Gln Ala Leu Leu Gly Asp Ala Arg Trp Leu Ala Ser Leu Val P $\overset{r}{ ext{ro}}$ Asn 155 Ser Ala Arg Gly Ala Thr Leu Arg Arg Leu Gly Ile Thr Asp Val Ala 170 165 Asp Leu Arg Ser Ser Arg Glu Val Ala Arg Arg Gly Pro Gly Arg Val Pro Asp Gly Ile Asp Val His Leu Leu Pro Phe Pro Asp Leu Ala Asp 200 Asp Asp Ala Asp Asp Ser Ala Pro His Glu Thr Ala Phe Lys Arg Leu 215 210 Leu Thr Asn Asp Gly Ser Asn Gly Glu Ser Gly Glu Ser Ser Gln Ser 235 230 Ile Asn Asp Ala Ala Thr Arg Tyr Met Thr Asp Glu Tyr Arg Gln Phe 250 . 245 Pro Thr Arg Asn Gly Ala Gln Arg Ala Leu His Arg Val Val Thr Leu 265 Leu Ala Ala Gly Arg Pro Val Leu Thr His Cys Phe Ala Gly Lys Asp 280 Arg Thr Gly Phe Val Val Ala Leu Val Leu Glu Ala Val Gly Leu Asp 300 295 290 Arg Asp Val Ile Val Ala Asp
- (2) INFORMATION FOR SEQ ID NO:184:
 - (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 2072 base pairs

(B) TYPE: nucleic acid(C) STRANDEDNESS: single(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:184:

CTCGTGCCGA	TTCGGCACGA	GCTGAGCAGC	CCAAGGGGCC	GTTCGGCGAA	GTCATCGAGG	60
CATTCGCCGA	CGGGCTGGCC	GGCAAGGGTA	AGCAAATCAA	CACCACGCTG	AACAGCCTGT	120
CGCAGGCGTT	GAACGCCTTG	AATGAGGGCC	GCGGCGACTT	CTTCGCGGTG	GTACGCAGCC	180
TGGCGCTATT	CGTCAACGCG	CTACATCAGG	ACGACCAAĊA	GTTCGTCGCG	TTGAACAAGA	240
ACCTTGCGGA	GTTCACCGAC	AGGTTGACCC	ACTCCGATGC	GGACCTGTCG	AACGCCATCC	300
AGCAATTCGA	CAGCTTGCTC	GCCGTCGCGC	GCCCGTTCTT	CGCCAAGAAC	CGCGAGGTGC	360
TGACGCATGA	CGTCAATAAT	CTCGCGACCG	TGACCACCAC	GTTGCTGCAG	CCCGATCCGT	420
TGGATGGGTT	GGAGACCGTC	CTGCACATCT	TCCCGACGCT	GGCGGCGAAC	ATTAACCAGC	480
TTTACCATCC	GACACACGGT	GGCGTGGTGT	CGCTTTCCGC	GTTCACGAAT	TTCGCCAACC	540
CGATGGAGTT	CATCTGCAGC	TCGATTCAGG	CGGGTAGCCG	GCTCGGTTAT	CAAGAGTCGG	600
CCGAACTCTG	TGCGCAGTAT	CTGGCGCCAG	TCCTCGATGC	GATCAAGTTC	AACTACTTTC	. 660
CGTTCGGCCT	GAACGTGGCC	AGCACCGCCT	CGACACTGCC	TAAAGAGATC	GCGTACTCCG	720
AGCCCCGCTT	GCAGCCGCCC	AACGGGTACA	AGGACACCAC	GGTGCCCGGC	ATCTGGGTGC	780
CGGATACGCC	GTTGTCACAC	CGCAACACGC	AGCCCGGTTG	GGTGGTGGCA	CCCGGGATGC	840
AAGGGGTTCA	GGTGGGACCG	ATCACGCAGG	GTTTGCTGAC	GCCGGAGTCC	CTGGCCGAAC	900
TCATGGGTGG	TCCCGATATC	GCCCCTCCGT	CGTCAGGGCT	GCAAACCCCG	CCCGGACCCC	960
CGAATGCGTA	CGACGAGTAC	CCCGTGCTGC	CGCCGATCGG	TTTACAGGCC	CCACAGGTGC	1020
CGATACCACC	GCCGCCTCCT	GGGCCCGACG	TAATCCCGGG	TCCGGTGCCA	CCGGTCTTGG	1080
CGGCGATCGT	GTTCCCAAGA	GATCGCCCGG	CAGCGTCGGA	AAACTTCGAC	TACATGGGCC	1140
TCTTGTTGCT	GTCGCCGGGC	CTGGCGACCT	TCCTGTTCGG	GGTGTCATCT	AGCCCCGCCC	1200
GTGGAACGAT	GGCCGATCGG	CACGTGTTGA	TACCGGCGAT	CACCGGCCTG	GCGTTGATCG	1260
CGGCATTCGT	CGCACATTCG	TGGTACCGCA	CAGAACATCC	GCTCATAGAC	ATGCGCTTGT	1320
TCCAGAACCG	AGCGGTCGCG	CAGGCCAACA	TGACGATGAC	GGTGCTCTCC	CTCGGGCTGT	1380

TTGGCTCCTT	CTTGCTGCTC	CCGAGCTACC	TCCAGCAAGT	GTTGCACCAA	TCACCGATGC	1440
AATCGGGGGT	GCATATCATC	CCACAGGGCC	TCGGTGCCAT	GCTGGCGATG	CCGATCGCCG	1500
GAGCGATGAT	GGACCGACGG	GGACCGGCCA	AGATCGTGCT	GGTTGGGATC	ATGCTGATCG	1560
CTGCGGGGTT	GGGCACCTTC	GCCTTTGGTG	TCGCGCGGCA	AGCGGACTAC	TTACCCATTC	1620
TGCCGACCGG	GCTGGCAATC	ATGGGCATGG	GCATGGGCTG	CTCCATGATG	CCACTGTCCG	1680
GGGCGGCAGT	GCAGACCCTG	GCCCCACATC	AGATCGCTCG	CGGTTCGACG	CTGATCAGCG	1740
TCAACCAGCA	GGTGGGCGGT	TCGATAGGGA	CCGCACTGAT	GTCGGTGCTG	CTCACCTACC	1800
AGTTCAATCA	CAGCGAAATC	ATCGCTACTG	CAAAGAAAGT	CGCACTGACC	CCAGAGAGTG	1860
GCGCCGGGCG	GGGGGCGCG	GTTGACCCTT	CCTCGCTACC	GCGCCAAACC	AACTTCGCGG	1920
CCCAACTGCT	GCATGACCTT	TCGCACGCCT	ACGCGGTGGT	ATTCGTGATA	GCGACCGCGC	1980
TAGTGGTCTC	GACGCTGATC	CCCGCGGCAT	TCCTGCCGAA	ACAGCAGGCT	AGTCATCGAA	2040
GAGCACCGTT	GCTATCCGCA	TGACGTCTGC	TT			2072

(2) INFORMATION FOR SEQ ID NO:185:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1923 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:185:

TCACCCCGGA	GAAGTCGTTC	GTCGACGACC	TGGACATCGA	CTCGCTGTCG	ATGGTCGAGA	60
TCGCCGTGCA	GACCGAGGAC	AAGTACGGCG	TCAAGATCCC	CGACGAGGAC	CTCGCCGGTC	120
TGCGTACCGT	CGGTGACGTT	GTCGCCTACA	TCCAGAAGCT	CGAGGAAGAA	AACCCGGAGG	180
CGGCTCAGGC	GTTGCGCGCG	AAGATTGAGT	CGGAGAACCC	CGATGCGGCA	CGAGCAGATC	240
GGTGCGTTTC	ACCCACATCG	CAAGCTCGAG	ACGCCCGTCG	TCCTCTTGCA	CGCTCAGCCA	300
GGTTGGCGTG	TCGCCGCCTT	CCAGCAAGTG	TTCCCACCAC	ACGAAGGGAC	CCTCGCGAAA	360
GGTGACTGAT	CCGCGGACCA	CATAGTCGAT	GCCACCGTGG	CTGACAATTG	CGCCGGGTCC	420
GAGTTGGCGG	GGGCCGAATT	GCGGCATTGC	GTCGAAGGCC	AGCGGATCCC	GGCGCCCGCC	480

PCT/US97/18214

CGGCGTGGCT	GGTGTTTTGG	GCCGCCGGAT	GGCCACGACG	AGAACGACGA	TGGCGGCGAT	540
GAACAGCGCC	ACGGCAATCA	CGACCAGCAG	ATTTCCCACG	CATACCCTCT	CGTACCGCTG	600
CGCCGCGGTT	GGTCGATCGG	TCGCATATCG	ATGGCGCCGT	TTAACGTAAC	AGCTTTCGCG	660
GGACCGGGGG	TCACAACGGG	CGAGTTGTCC	GGCCGGGAAC	CCGGCAGGTC	TCGGCCGCGG	720
TCACCCCAGC	TCACTGGTGC	ACCATCCGGG	TGTCGGTGAG	CGTGCAACTC	AAACACACTC	780
AACGGCAACG	GTTTCTCAGG	TCACCAGCTC	AACCTCGACC	CGCAATCGCT	CGTACGTTTC	840
GACCGCGCGC	AGGTCGCGAG	TCAGCAGCTT	TGCGCCGGCA	GCTTTCGCCG	TGAAGCCGAC	900
CAGGGCATCG	TAGGTTGCGC	CACCGGTGAC	ATCGTGCTCG	GCGAGGTGGT	CGGTCAAGCC	960
GCGATATGAG	CAGGCATCCA	GTGCCAGGTA	GTTGCTGGAG	GTGATGTCCG	CCAAGTAGGC	1020
GTGGACGGCA	ACAGGGGCAA	TACGATGCGG	CGGTGGTAGC	CGGGTCAAGA	CCGAATAGGT	1080
TTCCACAGCC	GCGTGCGCGA	TCAGATGGAC	GCCACGGTTG	AGCGCGCGCA	CGGCGGCCTC	1140
GTGCCCTTCG	TGCCAGGTCG	CGAATCCGGC	AACCAGCACG	CTGGTGTCTG	GTGCGATCAC	1200
CGCCGTGTGC	GATCGAGCGT	TTCCCGAACG	ATTTCGTCGG	TCAACGGGGG	CAGGGGACGT	1260
TCTGGCCGTG	CGACGAGAAC	CGAGCCTTCC	CGAACGAGTT	CGACACCGGT	CGGGGCCGGC	1320
TCAATCTCGA	TGCGCCCATC	GCGCTCGGTG	ATCTCCACCT	GGTCGTTCCC	GCGCAAGCCA	1380
AGGCGCTCGC	GAATCCGCTT	GGGAATCACC	AGACGTCCTG	CGACATCGAT	GGTTGTTCGC	1440
ATGGTAGGAA	ATTTACCATC	GCACGTTCCA	TAGGCGTGTC	CTGCGCGGGA	TGTCGGGACG	1500
ATCCGCTAGC	GTATCGAACG	ATTGTTTCGG	AAATGGCTGA	GGGAGCGTGC	GGTGCGGGTG	1560
ATGGGTGTCG	ATCCCGGGTT	GACCCGATGC	GGGCTGTCGC	TCATCGAGAG	TGGGCGTGGT	1620
CGGCAGCTCA	CCGCGCTGGA	TGTCGACGTG	GTGCGCACAC	CGTCGGATGC	GGCCTTGGCG	1680
CAGCGCCTGT	TGGCCATCAG	CGATGCCGTC	GAGCACTGGC	TGGACACCCA	TCATCCGGAG	1740
GTGGTGGCTA	TCGAACGGGT	GTTCTCTCAG	CTCAACGTGA	CCACGGTGAT	GGGCACCGCG	1800
CAGGCCGGCG	GCGTGATCGC	CCTGGCGGCG	GCCAAACGTG	GTGTCGACGT	GCATTTCCAT	1860
ACCCCCAGCG	AGGTCAAGGC	GGCGGTCACT	GGCAACGGTT	CCGCAGACAA	GGCTCAGGTC	1920
ACC						1923

- (2) INFORMATION FOR SEQ ID NO:186:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1055 base pairs
 - (B) TYPE: nucleic acid

- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:186:

CTGGCGTGCC	AGTGTCACCG	GCGATATGAC	GTCGGCATTC	AATTTCGCGG	CCCCGCCGGA	60
CCCGTCGCCA	CCCAATCTGG	ACCACCCGGT	CCGTCAATTG	CCGAAGGTCG	CCAAGTGCGT	120
GCCCAATGTG	GTGCTGGGTT	TCTTGAACGA	AGGCCTGCCG	TATCGGGTGC	CCTACCCCCA	180
AACAACGCCA	GTCCAGGAAT	CCGGTCCCGC	GCGGCCGATT	CCCAGCGGCA	TCTGCTAGCC	240
GGGGATGGTT	CAGACGTAAC	GGTTGGCTAG	GTCGAAACCC	GCGCCAGGGC	CGCTGGACGG	300
GCTCATGGCA	GCGAAATTAG	AAAACCCGGG	ATATTGTCCG	CGGATTGTCA	TACGATGCTG	360
AGTGCTTGGT	GGTTCGTGTT	TAGCCATTGA	GTGTGGATGT	GTTGAGACCC	TGGCCTGGAA	420
GGGGACAACG	TGCTTTTGCC	TCTTGGTCCG	CCTTTGCCGC	CCGACGCGGT	GGTGGCGAAA	480
CGGGCTGAGT	CGGGAATGCT	CGGCGGGTTG	TCGGTTCCGC	TCAGCTGGGG	AGTGGCTGTG	540
CCACCCGATG	ATTATGACCA	CTGGGCGCCT	GCGCCGGAGG	ACGGCGCCGA	TGTCGATGTC	600
CAGGCGGCCG	AAGGGGCGGA	CGCAGAGGCC	GCGGCCATGG	ACGAGTGGGA	TGAGTGGCAG	660
GCGTGGAACG	AGTGGGTGGC	GGAGAACGCT	GAACCCCGCT	TTGAGGTGCC	ACGGAGTAGC	720
AGCAGCGTGA	TTCCGCATTC	TCCGGCGGCC	GGCTAGGAGA	GGGGGCGCAG	ACTGTCGTTA	780
TTTGACCAGT	GATCGGCGGT	CTCGGTGTTC	CCGCGGCCGG	CTATGACAAC	AGTCAATGTG	840
CATGACAAGT	TACAGGTATT	AGGTCCAGGT	TCAACAAGGA	GACAGGCAAC	ATGGCAACAC	900
GTTTTATGAC	GGATCCGCAC	GCGATGCGGG	ACATGGCGGG	CCGTTTTGAG	GTGCACGCCC	960
AGACGGTGGA	GGACGAGGCT	CGCCGGATGT	GGGCGTCCGC	GCAAAACATC	TCGGGNGCGG	1020
GCTGGAGTGG	CATGGCCGAG	GCGACCTCGC	TAGAC			1055

(2) INFORMATION FOR SEQ ID NO:187:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 359 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUE	ENCE DESCRIPTION:	SEQ ID NO:18	7:		
CCGCCTCGTT GT	rggcatac tccgccgc	GG CCGCCTCGAC	CGCACTGGCC	GTGGCGTGTG	60
TCCGGGCTGA CCA	ACCGGGAT CGCCGAAC	CA TCCGAGATCA	CCTCGCAATG	ATCCACCTCG	120
CGCAGCTGGT CAC	CCCAGCCA CCGGGCGG	rg tgcgacagcg	CCTGCATCAC	CTTGGTATAG	180
CCGTCGCGCC CCA	AGCCGCAG GAAGTTGTA	AG TACTGGCCCA	CCACCTGGTT	ACCGGGACGG	240
GAGAAGTTCA GGG	STGAAGGT CGGCATGTC	CG CCGCCGAGGT	AGTTGACCCG	GAAAACCAGA	300
TCCTCCGGCA GGT	CCTCGGG CCCGCGCC	AC ACGACAAACC	CGACGCCGGG	ATAGGTCAG	359
(2) INFORMATIO	ON FOR SEQ ID NO:	188:			

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 350 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:188:

AACGGGCCCG TGGGCACCGC TCCTCTAAGG GCTCTCGTTG GTCGCATGAA GTGCTGGAAG 60
GATGCATCTT GGCAGATTCC CGCCAGAGCA AAACAGCCGC TAGTCCTAGT CCGAGTCGCC 120
CGCAAAGTTC CTCGAATAAC TCCGTACCCG GAGCGCCAAA CCGGGTCTCC TTCGCTAAGC 180
TGCGCGAACC ACTTGAGGTT CCGGGACTCC TTGACGTCCA GACCGATTCG TTCGAGTGGC 240
TGATCGGTTC GCCGCGTGG CGCGAATCCG CCGCCGAGCG GGGTGATGTC AACCCAGTGG 300
GTGGCCTGGA AGAGGTGCTC TACGAGCTGT CTCCGATCGA GGACTTCTCC 350

- (2) INFORMATION FOR SEQ ID NO:189:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 679 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:189:

Glu 1	Gln	Pro	Lys	Gly 5	Pro	Phe	Gly	Glu	Val 10	Ile	Glu	Ala	Phe	Ala 15	Asp
Gly	Leu	Ala	Gly 20	Lys	Gly	Lys	Gln	Ile 25	Asn	Thr	Thr	Leu	Asn 30	Ser	Leu
Ser	Ģln	Ala 35	Leu	Asn	Ala	Leu	Asn 40	Glu	Gly	Arg	Gly	Asp 45	Phe	Phe	Ala
Val	Val 50	Arg	Ser	Leu	Ala	Leu 55	Phe	Val	Asn	Ala	Leu 60	His	Gln	Asp	Asp
Gln 65	Gln.	Phe	Vаl	Ala	Leu 70	Asn	Lys	Asn	Leu	Ala 75	Glu	Phe	Thr	Asp	Arg 80
Leu	Thr	His	Ser	Asp 85	Ala	Asp	Leu	Ser	Asn 90	Ala	Ile	Gln	Gln	Phe 95	Asp
Ser	Leu	Leu	Ala 100	Val	Ala	Arg	Pro	Phe 105	Phe	Ala	Lys	Asn	Arg 110	Glu	Val
Leu	Thr	His 115	Asp	Val	Asn	Asn	Leu 120	Ala	Thr	Val	Thr	Thr 125	Thr	Leu	Leu
Gln	Pro 130	Asp	Pro	Leu	Asp	Gly 135	Leu	Glu	Thr	Val	Leu 140	His	Ile	Phe	Pro
Thr 145	Leu	Ala	Ala	Asn	Ile 150	Asn	Gln	Leu	Туr	His 155	Pro	Thr	His	Gly	Gly 160
Val	Val	Ser	Leu	Ser 165	Ala	Phe	Thr	Asn	Phe 170	Ala	Asn	Pro	Met	Glu 175	Phe
Ile	Cys	Ser	Ser 180	Ile	Gln	Ala	Gly	Ser 185	Arg	Leu	Gly	Tyr	Gln 190	Glu	Ser
Ala	Glu	Leu 195	Cys	Ala	Gln	Tyr	Leu 200	Ala	Pro	Val	Leu	Asp 205	Ala	Ile	Lys
Phe	Asn 210	Tyr	Phe	Pro	Phe	Gly 215		Asn	Val	Ala	Ser 220	Thr	Ala	Ser	Thr
Leu 225		Lys	Glu	Ile	Ala 230	Tyr	Ser	Glu	Pro	235	Let	Glr	Pro	Pro	240
Gly	Tyr	Lys	Asp	Thr 245		Val	Pro	Gly	11e 250	e Trp	Val	Pro	Asp	255	Pro
Leu	Ser	His	Arg 260		Thr	Gln	Pro	Gly 265	Trp	val	. Val	. Ala	270	Gly	/ Met
Gln	Gly	Val 275		Val	Gly	Pro	11e 280		Glr	n Gly	, Lei	1 Let 285	ı Thi	r Pro	o Glu

Ser	Leu 290	Ala	Glu	Leu	Met	Gly 295	Gly	Pro	Asp	Ile	Ala 300	Pro	Pro	Ser	Ser
Gly 305	Leu	Gln	Thr	Pro	Pro 310	Gly	Pro	Pro	Asn	Ala 315	Tyr	Asp	Glu	Tyr	Pro 320
Val	Leu	Pro	Pro	Ile 325	Gly	Leu	Gln	Ala	Pro 330	Gln	Val	Pro	Ile	Pro 335	Pro
Pro	Pro	Pro	Gly 340	Pro	Asp	Val	Ile	Pro 345	Gly	Pro	Val	Pro	Pro 350	Val	Leu
Ala	Ala	11e 355	Val	Phe	Pro	Arg	Asp 360	Arg	Pro	Ala	Ala	Ser 365	Glu	Asn	Phe
Asp	Tyr 370	Met	Gly	Leu	Leu	Leu 375	Leu	Ser	Pro	Gly	Leu 380	Ala	Thr	Phe	Leu
385					390					395	Met				400
				405					410		Ile			415	
			420					425			Ile		430		
		435					440				Thr	445			
	450					455					Pro 460				
465					470					475					480
				485					490		Ala			495	
			500					505			Gly		510		
		515					520				Ala	525			
_	530					535					Met 540				
545	_				550					555					560
				565					570	ı	: Ser			575	
Val	Gly	Gly	Ser	Ile	Gly	Thr	Ala	Leu	Met	Ser	Val	Leu	Leu	Thr	Tyr

580 585 590

Gln Phe Asn His Ser Glu Ile Ile Ala Thr Ala Lys Lys Val Ala Leu 595 600 605

Thr Pro Glu Ser Gly Ala Gly Arg Gly Ala Ala Val Asp Pro Ser Ser 610 615 620

Leu Pro Arg Gln Thr Asn Phe Ala Ala Gln Leu Leu His Asp Leu Ser 625 630 635 640

His Ala Tyr Ala Val Val Phe Val Ile Ala Thr Ala Leu Val Val Ser 645 650 655

Thr Leu Ile Pro Ala Ala Phe Leu Pro Lys Gln Gln Ala Ser His Arg 660 665 670

Arg Ala Pro Leu Leu Ser Ala 675

(2) INFORMATION FOR SEQ ID NO:190:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 120 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:190:

Thr Pro Glu Lys Ser Phe Val Asp Asp Leu Asp Ile Asp Ser Leu Ser

Met Val Glu Ile Ala Val Gln Thr Glu Asp Lys Tyr Gly Val Lys Ile 20 25 30

Pro Asp Glu Asp Leu Ala Gly Leu Arg Thr Val Gly Asp Val Val Ala 35 40 45

Tyr Ile Gln Lys Leu Glu Glu Glu Asn Pro Glu Ala Ala Gln Ala Leu 50 55 60

Arg Ala Lys Ile Glu Ser Glu Asn Pro Asp Ala Ala Arg Ala Asp Arg 65 70 75 80

Cys Val Ser Pro Thr Ser Gln Ala Arg Asp Ala Arg Arg Pro Leu Ala

Arg Ser Ala Arg Leu Ala Cys Arg Arg Leu Pro Ala Ser Val Pro Thr

Thr Arg Arg Asp Pro Arg Glu Arg 115 120

- (2) INFORMATION FOR SEQ ID NO:191:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 89 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:191:

Leu Ala Cys Gln Cys His Arg Arg Tyr Asp Val Gly Ile Gln Phe Arg 1 10 15

Gly Pro Ala Gly Pro Val Ala Thr Gln Ser Gly Pro Pro Gly Pro Ser 20 25 30

Ile Ala Glu Gly Arg Gln Val Arg Ala Gln Cys Gly Ala Gly Phe Leu $35 \hspace{1.5cm} 40 \hspace{1.5cm} 45$

Glu Arg Arg Pro Ala Val Ser Gly Ala Leu Pro Pro Asn Asn Ala Ser 50 55 60

Pro Gly Ile Arg Ser Arg Ala Ala Asp Ser Gln Arg His Leu Leu Ala 65 70 75 80

Gly Asp Gly Ser Asp Val Thr Val Gly 85

- (2) INFORMATION FOR SEQ ID NO:192:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 119 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:192:

Ala Ser Leu Leu Ala Tyr Ser Ala Ala Ala Ala Ser Thr Ala Leu Ala 1 5 10 15

Val Ala Cys Val Arg Ala Asp His Arg Asp Arg Arg Thr Ile Arg Asp 20 25 30

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His Leu Ala Met Ile His Leu Ala Gln Leu Val Thr Gln Pro Pro Gly 35 40 45

Gly Val Arg Gln Arg Leu His His Leu Gly Ile Ala Val Ala Pro Gln 50 55

Pro Gln Glu Val Val Leu Ala His His Leu Val Thr Gly Thr Gly 65 70 75 80

Glu Val Gln Gly Glu Gly Arg His Val Ala Ala Glu Val Val Asp Pro

Glu Asn Gln Ile Leu Arg Gln Val Leu Gly Pro Ala Pro His Asp Lys
100 105 110

Pro Asp Ala Gly Ile Gly Gln 115

(2) INFORMATION FOR SEQ ID NO:193:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 116 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:193:

Arg Ala Arg Gly His Arg Ser Ser Lys Gly Ser Arg Trp Ser His Glu
1 5 10 15

Val Leu Glu Gly Cys Ile Leu Ala Asp Ser Arg Gln Ser Lys Thr Ala 20 25 30

Ala Ser Pro Ser Pro Ser Arg Pro Gln Ser Ser Ser Asn Asn Ser Val

Pro Gly Ala Pro Asn Arg Val Ser Phe Ala Lys Leu Arg Glu Pro Leu

Glu Val Pro Gly Leu Leu Asp Val Gln Thr Asp Ser Phe Glu Trp Leu
65 70 75 80

Ile Gly Ser Pro Arg Trp Arg Glu Ser Ala Ala Glu Arg Gly Asp Val

Asn Pro Val Gly Gly Leu Glu Glu Val Leu Tyr Glu Leu Ser Pro Ile 100 105 110

Glu Asp Phe Ser 115

(2) INFORMATION FOR SEQ ID NO:194:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 811 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:194:

TGCTACGCAG	CAATCGCTTT	GGTGACAGAT	GTGGATGCCG	GCGTCGCTGC	TGGCGATGGC	60
GTGAAAGCCG	CCGACGTGTT	CGCCGCATTC	GGGGAGAACA	TCGAACTGCT	CAAAAGGCTG	120
GTGCGGGCCG	CCATCGATCG	GGTCGCCGAC	GAGCGCACGT	GCACGCACTG	TCAACACCAC	180
GCCGGTGTTC	CGTTGCCGTT	CGAGCTGCCA	TGAGGGTGCT	GCTGACCGGC	GCGGCCGGCT	240
TCATCGGGTC	GCGCGTGGAT	GCGGCGTTAC	GGGCTGCGGG	TCACGACGTG	GTGGGCGTCG	300
ACGCGCTGCT	GCCCGCCGCG	CACGGGCCAA	ACCCGGTGCT	GCCACCGGGC	TGCCAGCGGG	360
TCGACGTGCG	CGACGCCAGC	GCGCTGGCCC	CGTTGTTGGC	CGGTGTCGAT	CTGGTGTGTC	420
ACCAGGCCGC	CATGGTGGGT	. GCCGGCGTCA	ACGCCGCCGA	CGCACCCGCC	TATGGCGGCC	480
ACAACGATTT	CGCCACCACG	GTGCTGCTGG	CGCAGATGTT	CGCCGCCGGG	GTCCGCCGTT	540
TGGTGCTGGC	GTCGTCGATG	GTGGTTTACG	GGCAGGGGCG	CTATGACTGT	CCCCAGCATG	600
GACCGGTCGA	CCCGCTGCCG	CGGCGGCGAG	CCGACCTGGA	CAATGGGGTC	TTCGAGCACC	660
GTTGCCCGGG	GTGCGGCGAG	CCAGTCATCT	GGCAATTGGT	CGACGAAGAT	GCCCCGTTGC	720
GCCCGCGCAG	CCTGTACGCG	GCAGCAAGAC	CGCGCAGGAG	CACTACGCGC	TGGCGTGGTC	780
GGAAACGAAT	GGCGGTTCCG	TGGTGGCGTT	G			811

(2) INFORMATION FOR SEQ ID NO:195:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 966 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENC	E DESCRIPTION:	SEQ	ID	NO:195:
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GTCCCGCGAT	GTGGCCGAGC	ATGACTTTCG	GCAACACCGG	CGTAGTAGTC	GAAGATATCG	60
GACTTTGTGG	TCCCGGTGGC	GGGATAGAGC	ACCTGTCGGC	GTTGGTCAGC	GTCACCCGTT	120
GCTCGGACGC	CGAACCCATG	CTTTCAACGT	AGCCTGTCGG	TCACACAAGT	CGCGAGCGTA	180
ACGTCACGGT	CAAATATCGC	GTGGAATTTC	GCCGTGACGT	TCCGCTCGCG	GACAATCAAG	240
GCATACTCAC	TTACATGCGA	GCCATTTGGA	CGGGTTCGAT	CGCCTTCGGG	CTGGTGAACG	300
TGCCGGTCAA	GGTGTACAGC	GCTACCGCAG	ACCACGACAT	CAGGTTCCAC	CAGGTGCACG	360
CCAAGGACAA	CGGACGCATC	CGGTACAAGC	GCGTCTGCGA	GGCGTGTGGC	GAGGTGGTCG	420
ACTACCGCGA	TCTTGCCCGG	GCCTACGAGT	CCGGCGACGG	CCAAATGGTG	GCGATCACCG	480
ACGACGACAT	CGCCAGCTTG	CCTGAAGAAC	GCAGCCGGGA	GATCGAGGTG	TTGGAGTTCG	540
TCCCCGCCGC	CGACGTGGAC	CCGATGATGT	TCGACCGCAG	CTACTTTTTG	GAGCCTGATT	600
CGAAGTCGTC	GAAATCGTAT	GTGCTGCTGG	CTAAGACACT	CGCCGAGACC	GACCGGATGG	660
CGATCGTGGA	TCGCCCCACC	GGCCGTGAAT	GCAGGAAAAA	TAAGAGCCGC	TATCCACAAT	720
TCGGCGTCGA	GCTCGGCTAC	CACAAACGGT	AGAACGATCG	AGACATTCCC	GAGCTGAAGT	780
GCGGCGCTAT	AGAAGCCGCT	CTGCGCGATT	ATCAAACGCA	AAATACGCTT	ACTCATGCCA	840
TCGGCGCTGC	TCACCCGATG	CGACGTTTTT	GCCACGCTCC	ACCGCCTGCC	GCGCGACCTC	900
AAGTGGGCAT	GCATCCCACC	CGTTCCCGGA	AACCGGTTCC	GGCGGGTCGG	CTCATCGCTT	960
СВТССТ						966

(2) INFORMATION FOR SEQ ID NO:196:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 2367 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:196:

CCGCACCGCC	GGCAATACCG	CCAGCGCCAC	CGTTACCGCC	GTTTGCGCCG	TTGCCCCCGT	60
TGCCGCCCGT	cccgccggcc	CCGCCGATGG	AGTTCTCATC	GCCAAAAGTA	CTGGCGTTGC	120
CACCGGAGCC	GCCGTTGCCG	CCGTCACCGC	CAGCCCCGCC	GACTCCACCG	GCCCCACCGA	180

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CTCCGCCGCT	GCCACCGTTG	CCGCCGTTGC	CGATCAACAI	GCCGC1GGCG	CCACCCIIGC	240
CACCCACGCC	ACCGGCTCCG	CCCACCCCGC	CGACACCAAG	CGAGCTGCCG	CCGGAGCCAC	300
CATCACCACC	TACGCCACCG	ACCGCCCAGA	CACCAGCGAC	CGGGTCTTCG	TGAAACGTCG	360
CGGTGCCACC	ACCGCCGCCG	TTACCGCCAA	CCCCACCGGC	AACGCCGGCG	CCGCCATCCC	420
cecceeccc	GGCGTTGCCG	CCGTTGCCGC	CGTTGCCGAA	CAACAACCCG	CCGGCGCCGC	480
CGTTGCCGCC	CGCGCCGCCG	GTCCCGCCGG	CGCCGCCGAC	GCCAAGGCCG	CTGCCGCCCT	540
TGCCGCCATC	ACCACCCTTG	CCGCCGACCA	CATCGGGTTC	TGCCTCGGGG	TCTGGGCTGT	600
CAAACCTCGC	GATGCCAGCG	TTGCCGCCGC	TTCCCCCGGG	CCCCCCGTG	GCGCCGTCAC	660
CACCGATACC	ACCCGCGCCA	CCGGCGCCAC	CGTTGCCGCC	ATCACCGAAT	AGCAACCCGC	720
CGGCGCCACC	ATTGCCGCCA	GCTCCCCCTG	CGCCACCGTC	GGCGCCGGAG	GCGGCACTGG	780
CAGCCCCGTT	ACCACCGAAA	CCGCCGCTAC	CACCGGTAGA	GGTGGCAGTG	GCGATGTGTA	840
CGAAAGCGCC	GCCTCCGGCG	CCGCCGCTAC	CACCCCCACT	GCCGGCGGCT	ACACCGTCGG	900
ACCCGTTGCC	ACCATCACCG	CCAAAGGCGC	TCGCAATGTC	GCCCTGCGCG	ACTCCGCCGT	960
CGCCGCCGTT	GCCGCCGCCG	CCACCGGCAG	CGGCGGTACC	GCCGTCACCA	CCGGCACCGC	1020
CGGTGGCCTT	GCCCGAGCCT	GCCGTCGCGG	TGGCACCGTC	GCCGCCGGTG	CCACCGGTCG	1080
GCGTGCCGGC	AGTGCCATGG	CCGCCCGTGC	CGCCGTCGCC	GCCGGTTTGA	TCACCGATGC	1140
CGGACACATC	TGCCGGGCTG	TCCCCGGTGC	TGGCCGCGGG	GCCGGGCGTG	GGATTGACCC	1200
CGTTTGCCCC	GGCGAGGCCG	GCGCCGCCGG	TACCACCGGC	GCCGCCATGG	CCGAACAGCC	1260
CGGCGTTGCC	GCCGTTACCG	CCCGCACCCC	CGATGCCTGC	GGCCACGCTG	GTGCCGCCGA	1320
CACCGCCGTT	GCCGCCGTTG	CCCCACAACC	ACCCCCGTT	CCCACCGGCA	CCGCCGGCCG	1380
CGCCGGTACC	ACCGGCCCCG	CCGTTGCCGC	CGTTGCCGAT	CAACCCGGCC	GCGCCTCCGC	1440
TGCCGCCGGT	TTGACCGAAC	CCGCCAGCCG	CGCCGTTGCC	ACCGTTGCCA	AACAGCAACC	1500
CGCCGGCCGC	GCCAGGCTGC	CCGGGTGCCG	TCCCGTCGGC	GCCGTTTCCG	ATCAACGGGC	1560
GCCCCAAAAG	CGCCTCGGTG	GGCGCATTCA	CCGCACCCAG	CAGACTCCGC	TCAACAGCGG	1620
CTTCAGTGCT	GGCATACCGA	CCCGCGGCCG	CAGTCAACGC	CTGCACAAAC	TGCTCGTGAA	1680
ACGCTGCCAC	CTGTACGCTG	AGCGCCTGAT	ACTGCCGAGC	ATGGGCCCCG	AACAACCCCG	1740
CAATCCCCCC	CCACACTTCA	TCGGCAGCCG	CAGCCACCAC	TTCCGTCGTC	GGGATCGCCG	1800

CGGCCGCATT	AGCCGCGCTC	ACCTGCGAAC	CAATAGTCGA	TAAATCCAAA	GCCGCAGTTG	1860
CCAGCAGCTG	CGGCGTCGCG	ATCACCAAGG	ACACCTCGCA	CCTCCGGATA	CCCCATATCG	1920
CCGCACCGTG	TCCCCAGCGG	CCACGTGACC	TTTGGTCGCT	GGCTGGCGGC	CCTGACTATG	1980
GCCGCGACGG	CCCTCGTTCT	GATTCGCCCC	GGCGCGCAGC	TTGTTGCGCG	AGTTGAAGAC	2040
GGGAGGACAG	GCCGAGCTTG	GTGTAGACGT	GGGTCAAGTG	GGAATGCACG	GTCCGCGGCG	2100
AGATGAATAG	GCGGACGCCG	ATCTCCTTGT	TGCTGAGTCC	CTCACCGACC	AGTAGAGCCA	2160
CCTCAAGCTC	TGTCGGTGTC	AACGCGCCCC	AGCCACTTGT	CGGGCGTTTC	CGTGCACCGC	2220
GGCCTCGTTG	CGCGTACGCG	ATCGCCTCAT	CGATCGATAA	CGCAGTTCCT	TCGGCCCAGG	2280
CATCGTCGAA	CTCGCTGTCA	CCCATGGATT	TTCGAAGGGT	GGCTAGCGAC	GAGTTACAGC	2340
CCGCCTGGTA	GATCCCGAAG	CGGACCG			; ;	236

(2) INFORMATION FOR SEQ ID NO:197:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 376 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:197:

Gln Pro Ala Gly Ala Thr Ile Ala Ala Ser Ser Pro Cys Ala Thr Val 1 5 10 15

Gly Ala Gly Gly Gly Thr Gly Ser Pro Val Thr Thr Glu Thr Ala Ala 20 25 30

Thr Thr Gly Arg Gly Gly Ser Gly Asp Val Tyr Glu Ser Ala Ala Ser 35 40 45

Gly Ala Ala Ala Thr Thr Pro Thr Ala Gly Gly Tyr Thr Val Gly Pro 50 55 60

Val Ala Thr Ile Thr Ala Lys Gly Ala Arg Asn Val Ala Leu Arg Asp 65 70 75 80

Ser Ala Val Ala Ala Val Ala Ala Ala Ala Thr Gly Ser Gly Gly Thr 85 90 95.

Ala Val Thr Thr Gly Thr Ala Gly Gly Leu Ala Arg Ala Cys Arg Arg 100 105 110

Gly	Gly	Thr 115	Val	Ala	Ala	Gly	Ala 120	Thr	Gly	Arg	Arg	Ala 125	Gly	Ser	Ala
Met	Ala 130	Ala	Arg	Ala	Ala	Val 135	Ala	Ala	Gly	Leu	Ile 140	Thr	Asp	Ala	Gly
His 145	Ile	Cys	Arg	Ala	Val 150	Pro	Gly	Ala	Gly	Arg 155	Gly	Ala	Gly	Arg	Gly 160
Ile	Asp	Pro	Val	Cys 165	Pro	Gly	Glu	Ala	Gly 170	Ala	Ala	Gly	Thr	Thr 175	Gly
Ala	Ala	Met	Ala 180	Glu	Gln	Pro	Gly	Val 185	Ala	Ala	Val	Thr	Ala 190	Arg	Thr
Pro	Asp	Ala 195	Cys	Gly	His	Ala	Gly 200	Ala	Ala	Asp	Thr	Ala 205	Val	Ala /	Ala
Val	Ala 210	Pro	Gln	Pro	Pro	Pro 215	Val	Pro	Thr	Gly	Thr 220	Ala	Gly	Ar'g	Ala
Gly 225	Thr	Thr	Gly	Pro	Ala 230	Val	Ala	Ala	Val	Ala 235	Asp	Gln	Pro	Gly	Arg 240
Ala	Ser	Ala	Ala	Ala 245	Gly	Leu	Thr	Glu	Pro 250	Ala	Ser	Arg	Ala	Val 255	Ala
Thr	Val	Ala	Lys 260	Gln	Gln	Pro	Ala	Gly 265	Arg	Ala	Arg	Leu	Pro 270	Gly	Cys
Arg	Pro	Val 275	Gly	Ala	Val	Ser	Asp 280	Gln	Arg	Ala	Pro	Gln 285	Lys	Arg	Leu
Gly	Gly 290	Arg	Ile	His	Arg	Thr 295	Gln	Gln	Thr	Pro	Leu 300	Asn	Ser	Gly	Phe
Ser 305	Ala	Gly	Ile	Pro	Thr 310	Arg	Gly	Arg	Ser	Gln 315	Arg	Leu	His	Lys	Leu 320
Leu	Val	Lys	Arg	Cys 325	His	Leu	Tyr	Ala	Glu 330	Arg	Leu	Ile	Leu	Pro 335	Ser
Met	Gly	Pro	Glu 340	Gln	Pro	Arg	Asn	Arg 345	Arg	Arg	His	Phe	Ile 350	Gly	Ser
Arġ	Ser	His 355	His	Phe	Arg	Arg	Arg 360	Asp	Arg	Arg	Gly	Arg 365	Ile	Ser	Arg
Ala	His 370	Leu	Arg	Thr	Asn	Ser 375	Arg								

(2) INFORMATION FOR SEQ ID NO:198:

(i) SEQUENCE CHARACTERISTICS:

0818645A2 F 5

(A) LENGTH: 2852 base pairs

PCT/US97/18214

(B) TYPE: nucleic acid(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:198:

	CCCCGGCGAT					60
GACGTGGCGG	CGATGTTTGG	CTACCATGCC	GGGGCTTCGG	CGGCCGTCTC	GGCGTTGACA	120
CCGTTCGGCC	AGGCGCTGCC	GACCGTGGCG	GGCGGCGGTG	CGCTGGTCAG	CGCGGCCGCG	180
GCTCAGGTGA	CCACGCGGGT	CTTCCGCAAC	CTGGGCTTGG	CGAACGTCCG	ĊGAGGGCAĄĊ	240
GTCCGCAACG	GTAATGTCCG	GAACTTCAAT	CTCGGCTCGG	CCAACATCGG	CAACGGCAAC	300
ATCGGCAGCG	GCAACATCGG	CAGCTCCAAC	ATCGGGTTTG	GCAACGTGGG	TCCTGGGTTG	360
ACCGCAGCGC	TGAACAACAT	CGGTTTCGGC	AACACCGGÇA	GCAACAACAT	CGGGTTTGGC	420
AACACCGGCA	GCAACAACAT	CGGGTTCGGC	AATACCGGAG	ACGGCAACCG	AGGTATCGGG	480
CTCACGGGTA	GCGGTTTGTT	GGGGTTCGGC	GGCCTGAACT	CGGGCACCGG	CAACATCGGT	540
CTGTTCAACT	CGGGCACCGG	AAACGTCGGC	ATCGGCAACT	CGGGTACCGG	GAACTGGGGC	600
ATTGGCAACT	CGGGCAACAG	CTACAACACC	GGTTTTGGCA	ACTCCGGCGA	CGCCAACACG	660
GGCTTCTTCA	ACTCCGGAAT	AGCCAACACC	GGCGTCGGCA	ACGCCGGCAA	CTACAACACC	720
GGTAGCTACA	ACCCGGGCAA	CAGCAATACC	GGCGGCTTCA	ACATGGGCCA	GTACAACACG	780
GGCTACCTGA	ACAGCGGCAA	CTACAACACC	GGCTTGGCAA	ACTCCGGCAA	TGTCAACACC	840
GGCGCCTTCA	TTACTGGCAA	CTTCAACAAC	GGCTTCTTGT	GGCGCGGCGA	CCACCAAGGC	900
CTGATTTTCG	GGAGCCCCGG	CTTCTTCAAC	TCGACCAGTG	CGCCGTCGTC	GGGATTCTTC	960
AACAGCGGTG	CCGGTAGCGC	GTCCGGCTTC	CTGAACTCCG	GTGCCAACAA	TTCTGGCTTC	1020
TTCAACTCTT	CGTCGGGGGC	CATCGGTAAC	TCCGGCCTGG	CAAACGCGGG	GTGCTGGTA	1080
TCGGGCGTGA	TCAACTCGGG	CAACACCGTA	TCGGGTTTGT	TCAACATGAG	CCTGGTGGCC	1140
					GTCGGGATTT	1200
					GAACATTCTC	1260
					G TGACTTCAAC	1320
					A CGTCGGCAGC	1380
ATCCTIGGCE	7 00000111001					

TTCAATATCG GCAGTGGAAA CATCGGAGTA TTCAATGTCG GTTCCGGAAG CCTGGGAAAC	1440
TACAACATCG GATCCGGAAA CCTCGGGATC TACAACATCG GTTTTGGAAA CGTCGGCGAC	1500
TACAACGTCG GCTTCGGGAA CGCGGGCGAC TTCAACCAAG GCTTTGCCAA CACCGGCAAC	1560
AACAACATCG GGTTCGCCAA CACCGGCAAC AACAACATCG GCATCGGGCT GTCCGGCGAC	1620
AACCAGCAGG GCTTCAATAT TGCTAGCGGC TGGAACTCGG GCACCGGCAA CAGCGGCCTG	1680
TTCAATTCGG GCACCAATAA CGTTGGCATC TTCAACGCGG GCACCGGAAA CGTCGGCATC	1740
GCAAACTCGG GCACCGGGAA CTGGGGTATC GGGAACCCGG GTACCGACAA TACCGGCATC	1800
CTCAATGCTG GCAGCTACAA CACGGGCATC CTCAACGCCG GCGACTTCAA CACGGGCTTC	1860
TACAACACGG GCAGCTACAA CACCGGCGGC TTCAACGTCG GTAACACCAA CACCGGCAAC	1920
TTCAACGTGG GTGACACCAA TACCGGCAGC TATAACCCGG GTGACACCAA CACCGGCTTC	1980
TTCAATCCCG GCAACGTCAA TACCGGCGCT TTCGACACGG GCGACTTCAA CAATGGCTTC	2040
TTGGTGGCGG GCGATAACCA GGGCCAGATT GCCATCGATC TCTCGGTCAC CACTCCATTC	2100
ATCCCCATAA ACGAGCAGAT GGTCATTGAC GTACACAACG TAATGACCTT CGGCGGCAAC	2160
ATGATCACGG TCACCGAGGC CTCGACCGTT TTCCCCCAAA CCTTCTATCT GAGCGGTTTG	2220
TTCTTCTTCG GCCCGGTCAA TCTCAGCGCA TCCACGCTGA CCGTTCCGAC GATCACCCTC	2280
ACCATCGGCG GACCGACGGT GACCGTCCCC ATCAGCATTG TCGGTGCTCT GGAGAGCCGC	2340
ACGATTACCT TCCTCAAGAT CGATCCGGCG CCGGGCATCG GAAATTCGAC CACCAACCCC	2400
TCGTCCGGCT TCTTCAACTC GGGCACCGGT GGCACATCTG GCTTCCAAAA CGTCGGCGGC	2460
GGCAGTTCAG GCGTCTGGAA CAGTGGTTTG AGCAGCGCGA TAGGGAATTC GGGTTTCCAG	2520
AACCTCGGCT CGCTGCAGTC AGGCTGGGCG AACCTGGGCA ACTCCGTATC GGGCTTTTTC	2580
AACACCAGTA CGGTGAACCT CTCCACGCCG GCCAATGTCT CGGGCCTGAA CAACATCGGC	2640
ACCAACCTGT CCGGCGTGTT CCGCGGTCCG ACCGGGACGA TTTTCAACGC GGGCCTTGCC	2700
AACCTGGGCC AGTTGAACAT CGGCAGCGCC TCGTGCCGAA TTCGGCACGA GTTAGATACG	2760
GTTTCAACAA TCATATCCGC GTTTTGCGGC AGTGCATCAG ACGAATCGAA CCCGGGAAGC	2820
GTAAGCGAAT AAACCGAATG GCGGCCTGTC AT	2852

(2) INFORMATION FOR SEQ ID NO:199:

(i) SEQUENCE CHARACTERISTICS:

⁽A) LENGTH: 943 amino acids

- (B) TYPE: amino acid
- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:199:

Gly Gln Asn Ala Pro Ala Ile Ala Ala Thr Glu Ala Ala Tyr Asp Gln 1 5 10 15

Met Trp Ala Gln Asp Val Ala Ala Met Phe Gly Tyr His Ala Gly Ala 20 25 30

Ser Ala Ala Val Ser Ala Leu Thr Pro Phe Gly Gln Ala Leu Pro Thr 35 40 45

Val Ala Gly Gly Gly Ala Leu Val Ser Ala Ala Ala Ala Gln Val Thr 50 55 60

Thr Arg Val Phe Arg Asn Leu Gly Leu Ala Asn Val Arg Glu Gly Asn 65 70 75 80

Val Arg Asn Gly Asn Val Arg Asn Phe Asn Leu Gly Ser Ala Asn Ile 85 90 95

Gly Asn Gly Asn Ile Gly Ser Gly Asn Ile Gly Ser Ser Asn Ile Gly
100 105 110

Phe Gly Asn Val Gly Pro Gly Leu Thr Ala Ala Leu Asn Asn Ile Gly
115 120 125

Phe Gly Asn Thr Gly Ser Asn Asn Ile Gly Phe Gly Asn Thr Gly Ser 130 135 140

Asn Asn Ile Gly Phe Gly Asn Thr Gly Asp Gly Asn Arg Gly Ile Gly 145 150 155 160

Leu Thr Gly Ser Gly Leu Leu Gly Phe Gly Gly Leu Asn Ser Gly Thr 165 170 175

Gly Asn Ile Gly Leu Phe Asn Ser Gly Thr Gly Asn Val Gly Ile Gly 180 185 190

Asn Ser Gly Thr Gly Asn Trp Gly Ile Gly Asn Ser Gly Asn Ser Tyr 195 200 205

Asn Thr Gly Phe Gly Asn Ser Gly Asp Ala Asn Thr Gly Phe Phe Asn 210 215 220

Ser Gly Ile Ala Asn Thr Gly Val Gly Asn Ala Gly Asn Tyr Asn Thr 225 230 235 240

Gly	Ser	Tyr	Asn	Pro 245	Gly	Asn	Ser	Asn	Thr 250	Gly	Gly	Phe	Asn	Met 255	Gly
Gln	Tyr	Asn	Thr 260	Gly	Tyr	Leu	Asn	Ser 265	Gly	Asn	Tyr	Asn	Thr 270	Gly	Leu
Ala	Asn	Ser 275	Gly	Asn	Val	Asn	Thr 280	Gly	Ala	Phe	Ile	Thr 285	Gly	Asn	Phe
Asn	Asn 290	Gly	Phe	Leu	Trp	Arg 295	Gly	Asp	His	Gln	Gly 300	Leu	Ile	Phe	Gly
Ser 305	Pro	Gly	Phe	Phe	Asn 310	Ser	Thr	Ser	Ala	Pro 315	Ser	Ser	Gly	Phe	Phe 320
Asn	Ser	Gly	Ala	Gly 325	Ser	Ala	Ser	Gly	Phe 330	Leu	Asn	Ser	Gly	Ala 335	Asn
Asn	Ser	Gly	Phe 340	Phe	Asn	Ser	Ser	Ser 345	Gly	Ala	Ile	Gly	Asn 350	Ser	Gly
Leu	Ala	Asn 355	Ala	Gly	Val	Leu	Val 360	Ser	Gly	Val	Ile	Asn 365	Ser	Gly	Asn
Thr	Val 370	Ser	Gly	Leu	Phe	Asn 375	Met	Ser	Leu	Val	Ala 380	Ile	Thr	Thr	Pro
Ala 385	Leu	Ile	Ser	Gly	Phe 390	Phe	Asn	Thr	Gly	Ser 395	Asn	Met	Ser	Gly	Phe 400
Phe	Gly	Gly	Pro	Pro 405	Val	Phe	Asn	Leu	Gly 410	Leu	Ala	Asn	Arg	Gly 415	Val
Val	Asn	Ile	Leu 420		Asn	Ala	Asn	11e 425	Gly	Asn	Tyr	Asn	11e	Leu	Gly
Ser	Gly	Asn 435		Gly	Asp	Phe	Asn 440	Ile	Lev	ı Gly	ser,	Gly 445	Asr	Leu	Gly
Ser	Gln 450		Ile	Leu	Gly	Ser 455	Gly	Asn	val	L Gly	y Ser 460	Phe	e Asr	ı Ile	e Gly
Ser 465		Asn	Ile	Gly	Val 470		Asn	Val	Gly	y Sei 475	c Gly	, Ser	: Le	ı Gly	480
Tyr	Asn	Ile	Gly	Ser 485		Asn	Leu	Gl	/ Ile	e Ту: О	r Asr	ı Ile	e Gl	y Phe 499	e Gly
Asn	. Val	. Gly	Asp 500		Asn	Val	. Gly	Phe 505	e Gl	y Ası	n Ala	a Gly	y Ası 51	p Phe	e Asn
Glr	Gly	Phe 515		a Asn	Thr	Gly	7 Asr 520		n As	n Il	e Gl	y Pho 52	e Al	a Ası	n Thr
Gly	, Asr	Asr	n Asr	ıle	e Gly	, Ile	e Gly	, Le	ı Se	r Gl	y Ası	o As	n Gl	n Gl	n Gly

	530					535					540				
Phe 545	Asn	Ile	Ala	Ser	Gly 550	Trp	Asn	Ser	Gly	Thr 555	Gly	Asn	Ser	Gly	Leu 560
Phe	Asn	Ser	Gly	Thr 565	Asn	Asn	Val	Gly	Ile 570	Phe	Asn	Ala	Gly	Thr 575	Gly
Asn	Val	Gly	Ile 580	Ala	Asn	Ser	Gly	Thr 585	Gly	Asn	Trp	Gly	Ile 590	Gly	Asn
Pro	Gly	Thr 595	Asp	Asn	Thr	Gly	Ile 600	Leu	Asn	Ala	Gly	Ser 605	Tyr	Asn	Thr
Gly	Ile 610	Leu	Asn	Ala	Gly	Asp 615	Phe	Asn	Thr	Gly	Phe 620	Tyr	Asn	Thr	Gly
Ser 625	Tyr	Asn	Thr	Gly	Gly 630	Phe	Asn	Val	Gly	Asn 635	Thr	Asn	Thr	GÍY	Asn 640
Phe	Asn	Val	Gly	Asp 645	Thr	Asn	Thr	Gly	Ser 650	Tyr	Asn	Pro	Gly	Asp 655	Thr
Asn	Thr	Gly	Phe 660	Phe	Asn	Pro	Gly	Asn 665	Val	Asn	Thr	Gly	Ala 670	Phe	Asp
Thr	Gly	Asp 675	Phe	Asn	Asn	Gly	Phe 680	Leu	Val	Ala	Gly	Asp 685	Asn	Gln	Gly
Gln	Ile 690	Ala	Ile	Asp	Leu	Ser 695	Val	Thr	Thr	Pro	Phe 700	Ile	Pro	Ile	Asn
Glu 705	Gln	Met	Val	Ile	Asp 710	Val	His	Asn	Val	Met 715	Thr	Phe	Gly	Gly	Asn 720
Met	Ile	Thr	Val	Thr 725	Glu	Ala	Ser	Thr	Val 730	Phe	Pro	Gln	Thr	Phe 735	Tyr
Leu	Ser	Gly	Leu 740	Phe	Phe	Phe	Gly	Pro 745	Val	Asn	Leu	Ser	Ala 750	Ser	Thr
Leu	Thr	Val 755	Pro	Thr	Ile	Thr	Leu 760	Thr	Ile	Gly	Gly	Pro 765	Thr	Val	Thr
Val	Pro 770	Ile	Ser	Ile	Val	Gly 775	Ala	Leu	Glu	Ser	780	Thr	Ile	Thr	Phe
Leu 785	Lys	Ile	Asp	Pro	Ala 790	Pro	Gly	Ile	Gly	Asn 795	Ser	Thr	Thr	: Asr	800
Ser	Ser	Gly	Phe	Phe 805	Asn	Ser	Gly	Thr	Gly 810	Gly	Thr	Ser	: Gly	/ Phe 815	e Gln
Asn	Val	Gly	Gly 820	Gly	Ser	Ser	Gly	Val 825	Trp	Asr	Ser	Gly	/ Let 830	ı Sei	s Ser

Ala	Ile	Gly 835	Asn	Ser	Gly	Phe	Gln 840	Asn	Leu	Gly	Ser	Leu 845	Gln	Ser	Gly
Trp	Ala 850	Asn	Leu	Gly	Asn	Ser 855	Val	Ser	Gly	Phe	Phe 860	Asn	Thr	Ser	Th:
Val 865	Asn	Leu	Ser	Thr	Pro 870	Ala	Asn	Val	Ser	Gly 875	Leu	Asn	Asn	Ile	G1;
Thr	Asn	Leu	Ser	Gly 885	Val	Phe	Arg	Gly	Pro 890	Thr	Gly	Thr	Ile	Phe 895	As
Ala	Gly	Leu	Ala 900	Asn	Leu	Gly	Gln	Leu 905	Asn	Ile	Gly	Ser	Ala 910	Ser	Су
Arg	Ile	Arg 915	His	Glu	Leu	Asp	Thr 920	Val	Ser	Thr	Ile	Ile 925	Ser	Ala	Ph
Cys	Gly	Ser	Ala	Ser	Asp	Glu	Ser	Asn	Pro	Gly	Ser	Val	Ser	Glu	

(2) INFORMATION FOR SEQ ID NO:200:

930

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 53 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:200:

GGATCCATAT GGGCCATCAT CATCATCATC ACGTGATCGA CATCATCGGG ACC

935

53

- (2) INFORMATION FOR SEQ ID NO:201:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 42 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:201:

CCTGAATTCA GGCCTCGGTT GCGCCGGCCT CATCTTGAAC GA

(2) INFORMATION FOR SEQ ID NO:202:

....

 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 31 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(xi) SEQUENCE DESCRIPTION: SEQ ID 10:202:	
GGATCCTGCA GGCTCGAAAC CACCGAGCGG T	31
(2) INFORMATION FOR SEQ ID NO:203:	[
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 31 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	,
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:203:	
CTCTGAATTC AGCGCTGGAA ATCGTCGCGA T	31
(2) INFORMATION FOR SEQ ID NO:204:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 33 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:204:	
GGATCCAGCG CTGAGATGAA GACCGATGCC GCT	33
(2) INFORMATION FOR SEQ ID NO: 205:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 38 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	

60

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:205:	
GGATATCTGC AGAATTCAGG TTTAAAGCCC ATTTGCGA	38
(2) INFORMATION FOR SEQ ID NO:206:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 30 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:206:	٠.
CCGCATGCGA GCCACGTGCC CACAACGGCC	30
(2) INFORMATION FOR SEQ ID NO:207:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 37 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:207:	·
CTTCATGGAA TTCTCAGGCC GGTAAGGTCC GCTGCGG	37
(2) INFORMATION FOR SEQ ID NO:208:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 7676 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear 	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:208:	

TGGCGAATGG GACGCGCCCT GTAGCGGCGC ATTAAGCGCG GCGGGTGTGG TGGTTACGCG

CAGCGTGACC	GCTACACTTG	CCAGCGCCCT	AGCGCCCGCT	CCTTTCGCTT	TCTTCCCTTC	120
CTTTCTCGCC	ACGTTCGCCG	GCTTTCCCCG	TCAAGCTCTA	AATCGGGGGC	TCCCTTTAGG	180
GTTCCGATTT	AGTGCTTTAC	GGCACCTCGA	CCCCAAAAAA	CTTGATTAGG	GTGATGGTTC	240
ACGTAGTGGG	CCATCGCCCT	GATAGACGGT	TTTTCGCCCT	TTGACGTTGG	AGTCCACGTT	300
CTTTAATAGT	GGACTCTTGT	TCCAAACTGG	AACAACACTC	AACCCTATCT	CGGTCTATTC	360
TTTTGATTTA	TAAGGGATTT	TGCCGATTTC	GGCCTATTGG	TTAAAAAATG	AGCTGATTTA	420
ACAAAAATTT	AACGCGAATT	TTAACAAAAT	ATTAACGTTT	ACAATTTCAG	GTGGCACTTT	480
TCGGGGAAAT	GTGCGCGGAA	CCCCTATTTG	TTTATTTTTC	TAAATACATT	CAAATATGTA	540
TCCGCTCATG	AATTAATTCT	TAGAAAAACT	CATCGAGCAT	CAAATGAAAC	TGCAATTTAT	600
TCATATCAGG	ATTATCAATA	CCATATTTTT	GAAAAAGCCG	TTTCTGTAAT	GAAGGAGAAA	660
ACTCACCGAG	GCAGTTCCAT	AGGATGGCAA	GATCCTGGTA	TCGGTCTGCG	ATTCCGACTC	720
GTCCAACATC	AATACAACCT	ATTAATTTCC	CCTCGTCAAA	AATAAGGTTA	TCAAGTGAGA	780
AATCACCATG	AGTGACGACT	GAATCCGGTG	AGAATGGCAA	AAGTTTATGC	ATTTCTTTCC	840
AGACTTGTTC	AACAGGCCAG	CCATTACGCT	CGTCATCAAA	ATCACTCGCA	TCAACCAAAC	900
CGTTATTCAT	TCGTGATTGC	GCCTGAGCGA	GACGAAATAC	GCGATCGCTG	TTAAAAGGAC	960
AATTACAAAC	AGGAATCGAA	TGCAACCGGC	GCAGGAACAC	TGCCAGCGCA	TCAACAATAT	1020
TTTCACCTGA	ATCAGGATAT	TCTTCTAATA	CCTGGAATGC	TGTTTTCCCG	GGGATCGCAG	1080
TGGTGAGTAA	CCATGCATCA	TCAGGAGTAC	GGATAAAATG	CTTGATGGTC	GGAAGAGGCA	1140
TAAATTCCGT	CAGCCAGTTT	AGTCTGACCA	TCTCATCTGT	AACATCATTG	GCAACGCTAC	1200
CTTTGCCATG	TTTCAGAAAC	AACTCTGGCG	CATCGGGCTT	CCCATACAAT	CGATAGATTG	1260
TCGCACCTGA	TTGCCCGACA	TTATCGCGAG	CCCATTTATA	CCCATATAAA	TCAGCATCCA	1320
TGTTGGAATT	TAATCGCGGC	CTAGAGCAAG	ACGTTTCCCG	TTGAATATGG	CTCATAACAC	1380
CCCTTGTATT	ACTGTTTATG	TAAGCAGACA	GTTTTATTGT	TCATGACCAA	AATCCCTTAA	1440
CGTGAGTTTT	CGTTCCACTG	AGCGTCAGAC	CCCGTAGAAA	AGATCAAAGG	ATCTTCTTGA	1500
GATCCTTTTT	TTCTGCGCGT	AATCTGCTGC	TTGCAAACAA	AAAAACCAC	GCTACCAGCG	1560
GTGGTTTGTT	TGCCGGATCA	AGAGCTACCA	ACTCTTTTTC	CGAAGGTAAC	TGGCTTCAGC	162
7 C 7 C C C C 7 C 7	тассадатас	тстссттста	GTGTAGCCGT	AGTTAGGCCA	CCACTTCAAG	168

AACTCTGTAG	CACCGCCTAC	ATACCTCGCT	CTGCTAATCC	TGTTACCAGT	GGCTGCTGCC	1/40
AGTGGCGATA	AGTCGTGTCT	TACCGGGTTG	GACTCAAGAC	GATAGTTACC	GGATAAGGCG	1800
CAGCGGTCGG	GCTGAACGGG	GGGTTCGTGC	ACACAGCCCA	GCTTGGAGCG	AACGACCTAC	1860
ACCGAACTGA	GATACCTACA	GCGTGAGCTA	TGAGAAAGCG	CCACGCTTCC	CGAAGGGAGA	1920
AAGGCGGACA	GGTATCCGGT	AAGCGGCAGG	GTCGGAACAG	GAGAGCGCAC	GAGGGAGCTT	1980
CCAGGGGGAA	ACGCCTGGTA	TCTTTATAGT	CCTGTCGGGT	TTCGCCACCT	CTGACTTGAG	2040
CGTCGATTTT	TGTGATGCTC	GTCAGGGGGG	CGGAGCCTAT	GGAAAAACGC	CAGCAACGCG	2100
GCCTTTTTAC	GGTTCCTGGC	CTTTTGCTGG	CCTTTTGCTC	ACATGTTCTT	TCCTGCGTTA	2160
TCCCCTGATT	CTGTGGATAA	CCGTATTACC	GCCTTTGAGT	GAGCTGATAC	CGCTCGCCGC	2220
AGCCGAACGA	CCGAGCGCAG	CGAGTCAGTG	AGCGAGGAAG	CGGAAGAGCG	CCTGATGCGG	2280
TATTTTCTCC	TTACGCATCT	GTGCGGTATT	TCACACCGCA	TATATGGTGC	ACTCTCAGTA	2340
CAATCTGCTC	TGATGCCGCA	TAGTTAAGCC	AGTATACACT	CCGCTATCGC	TACGTGACTG	2400
GGTCATGGCT	GCGCCCGAC	ACCCGCCAAC	ACCCGCTGAC	GCGCCCTGAC	GGGCTTGTCT	2460
GCTCCCGGCA	TCCGCTTACA	GACAAGCTGT	GACCGTCTCC	GGGAGCTGCA	TGTGTCAGAG	2520
GTTTTCACCG	TCATCACCGA	AACGCGCGAG	GCAGCTGCGG	TAAAGCTCAT	CAGCGTGGTC	2580
GTGAAGCGAT	TCACAGATGT	CTGCCTGTTC	ATCCGCGTCC	AGCTCGTTGA	GTTTCTCCAG	2640
AAGCGTTAAT	GTCTGGCTTC	TGATAAAGCG	GGCCATGTTA	AGGGCGGTTT	TTTCCTGTTT	2700
GGTCACTGAT	GCCTCCGTGT	AAGGGGGATT	TCTGTTCATG	GGGGTAATGA	TACCGATGAA	2760
ACGAGAGAGG	ATGCTCACGA	TACGGGTTAC	TGATGATGAA	CATGCCCGGI	TACTGGAACG	2820
TTGTGAGGGT	AAACAACTGG	CGGTATGGAT	GCGGCGGGAC	CAGAGAAAA	TCACTCAGGG	2880
TCAATGCCAG	CGCTTCGTTA	ATACAGATGT	AGGTGTTCCA	CAGGGTAGCO	AGCAGCATCC	2940
TGCGATGCAG	ATCCGGAACA	TAATGGTGCA	GGGCGCTGAC	TTCCGCGTT	CCAGACTTTA	3000
CGAAACACGG	AAACCGAAGA	CCATTCATGT	TGTTGCTCAG	GTCGCAGACO	G TTTTGCAGCA	3060
GCAGTCGCTT	CACGTTCGCT	CGCGTATCGG	TGATTCATTC	TGCTAACCAG	TAAGGCAACC	3120
CCGCCAGCCT	AGCCGGGTCC	TCAACGACAG	GAGCACGATO	ATGCGCACCO	C GTGGGGCCGC	3180
CATGCCGGCG	ATAATGGCCT	GCTTCTCGCC	GAAACGTTTC	GTGGCGGGA	CAGTGACGAA	3240
GGCTTGAGCG	AGGGCGTGCA	AGATTCCGAA	TACCGCAAGO	GACAGGCCG	A TCATCGTCGC	3300
GCTCCAGCGA	AAGCGGTCCT	CGCCGAAAAT	GACCCAGAGO	C GCTGCCGGC	A CCTGTCCTAC	3360

WO 98/16645

PCT/US97/18214

GAGTTGCATG	ATAAAGAAGA	CAGTCATAAG	TGCGGCGACG	ATAGTCATGC	CCCGCGCCCA	3420
CCGGAAGGAG	CTGACTGGGT	TGAAGGCTCT	CAAGGGCATC	GGTCGAGATC	CCGGTGCCTA	3480
ATGAGTGAGC	TAACTTACAT	TAATTGCGTT	GCGCTCACTG	CCCGCTTTCC	AGTCGGGAAA	3540
CCTGTCGTGC	CAGCTGCATT	AATGAATCGG	CCAACGCGCG	GGGAGAGGCG	GTTTGCGTAT	3600
rgggcgccag	GGTGGTTTTT	CTTTTCACCA	GTGAGACGGG	CAACAGCTGA	TTGCCCTTCA	3660
CCGCCTGGCC	CTGAGAGAGT	TGCAGCAAGC	GGTCCACGCT	GGTTTGCCCC	AGCAGGCGAA	3720
AATCCTGTTT	GATGGTGGTT	AACGGCGGGA	TATAACATGA	GCTGTCTTCG	GTATCGTCGT	378.0
ATCCCACTAC	CGAGATATCC	GCACCAACGC	GCAGCCCGGA	CTCGGTAATG	GCGCGCATTG	3840
CGCCCAGCGC	CATCTGATCG	TTGGCAACCA	GCATCGCAGT	GGGAACGATG	CCCTCATTCA	3900
GCATTTGCAT	GGTTTGTTGA	AAACCGGACA	TGGCACTCCA	GTCGCCTTCC	CGTTCCGCTA	3960
TCGGCTGAAT	TTGATTGCGA	GTGAGATATT	TATGCCAGCC	AGCCAGACGC	AGACGCGCCG	4020
AGACAGAACT	TAATGGGCCC	GCTAACAGCG	CGATTTGCTG	GTGACCCAAT	GCGACCAGAT	4080
GCTCCACGCC	CAGTCGCGTA	CCGTCTTCAT	GGGAGAAAAT	AATACTGTTG	ATGGGTGTCT	4140
GGTCAGAGAC	ATCAAGAAAT	AACGCCGGAA	CATTAGTGCA	GGCAGCTTCC	ACAGCAATGG	4200
CATCCTGGTC	ATCCAGCGGA	TAGTTAATGA	TCAGCCCACT	GACGCGTTGC	GCGAGAAGAT	4260
TGTGCACCGC	CGCTTTACAG	GCTTCGACGC	CGCTTCGTTC	TACCATCGAC	ACCACCACGC	4320
TGGCACCCAG	TTGATCGGCG	CGAGATTTAA	TCGCCGCGAC	AATTTGCGAC	GGCGCGTGCA	4380
GGGCCAGACT	GGAGGTGGCA	ACGCCAATCA	GCAACGACTG	TTTGCCCGCC	AGTTGTTGTG	4440
CCACGCGGTT	GGGAATGTAA	TTCAGCTCCG	CCATCGCCGC	TTCCACTTTT	TCCCGCGTTT	4500
TCGCAGAAAC	GTGGCTGGCC	TGGTTCACCA	CGCGGGAAAC	GGTCTGATAA	GAGACACCGG	4560
CATACTCTGC	GACATCGTAT	AACGTTACTG	GTTTCACATT	CACCACCCTG	AATTGACTCT	462
CTTCCGGGCG	CTATCATGCC	ATACCGCGAA	AGGTTTTGCG	CCATTCGATG	GTGTCCGGGA	468
TCTCGACGCT	CTCCCTTATG	CGACTCCTGC	ATTAGGAAGC	AGCCCAGTAG	TAGGTTGAGG	474
CCGTTGAGCA	CCGCCGCCGC	AAGGAATGGT	GCATGCAAGG	AGATGGCGCC	CAACAGTCCC	480
CCGGCCACGG	GGCCTGCCAC	CATACCCACG	CCGAAACAAG	CGCTCATGAG	CCCGAAGTGG	486
CGAGCCCGAT	CTTCCCCATC	GGTGATGTCG	GCGATATAGG	CGCCAGCAAC	CGCACCTGTG	492
COCCCCCCC	TCCCGCCAC	GATGCGTCCG	GCGTAGAGGA	TCGAGATCTC	GATCCCGCGA	498

AATTAATACG	ACTCACTATA	GGGGAATTGT	GAGCGGATAA	CAATTCCCCT	CTAGAAATAA	5040
TTTTGTTTAA	CTTTAAGAAG	GAGATATACA	TATGGGCCAT	CATCATCATC	ATCACGTGAT	5100
CGACATCATC	GGGACCAGCC	CCACATCCTG	GGAACAGGCG	GCGGCGGAGG	CGGTCCAGCG	5160
GGCGCGGGAT	AGCGTCGATG	ACATCCGCGT	CGCTCGGGTC	ATTGAGCAGG	ACATGGCCGT	5220
GGACAGCGCC	GGCAAGATCA	CCTACCGCAT	CAAGCTCGAA	GTGTCGTTCA	AGATGAGGCC	5280
GGCGCAACCG	AGGGGCTCGA	AACCACCGAG	CGGTTCGCCT	GAAACGGGCG	CCGGCGCCGG	5340
TACTGTCGCG	ACTACCCCCG	CGTCGTCGCC	GGTGACGTTG	GCGGAGACCG	GTAGCACGCT	5400
GCTCTACCCG	CTGTTCAACC	TGTGGGGTCC	GGCCTTTCAC	GAGAGGTATC	CGAACGTCAC	5460
GATCACCGCT	CAGGGCACCG	GTTCTGGTGC	CGGGATCGCG	CAGGCCGCCG	CCGGGACGGT	5520
CAACATTGGG	GCCTCCGACG	CCTATCTGTC	GGAAGGTGAT	ATGGCCGCGC	ACAAGGGGCT	5580
GATGAACATC	GCGCTAGCCA	TCTCCGCTCA	GCAGGTCAAC	TACAACCTGC	CCGGAGTGAG	5640
CGAGCACCTC	AAGCTGAACG	GAAAAGTCCT	GGCGGCCATG	TACCAGGGCA	CCATCAAAAC	5700
CTGGGACGAC	CCGCAGATCG	CTGCGCTCAA	CCCCGGCGTG	AACCTGCCCG	GCACCGCGGT	5760
AGTTCCGCTG	CACCGCTCCG	ACGGGTCCGG	TGACACCTTC	TTGTTCACCC	AGTACCTGTC	5820
CAAGCAAGAT	CCCGAGGGCT	GGGGCAAGTC	GCCCGGCTTC	GGCACCACCG	TCGACTTCCC	5880
GGCGGTGCCG	GGTGCGCTGG	GTGAGAACGG	CAACGGCGGC	ATGGŤGACCG	GTTGCGCCGA	5940
GACACCGGGC	TGCGTGGCCT	ATATCGGCAT	CAGCTTCCTC	GACCAGGCCA	GTCAACGGGG	6000
ACTCGGCGAG	GCCCAACTAG	GCAATAGCTC	TGGCAATTTC	TTGTTGCCCG	ACGCGCAAAG	6060
CATTCAGGCC	GCGGCGGCTG	GCTTCGCATC	GAAAACCCCG	GCGAACCAGG	CGATTTCGAT	6120
GATCGACGGG	CCCGCCCCGG	ACGGCTACCC	GATCATCAAC	TACGAGTACG	CCATCGTCAA	6180
CAACCGGCAA	AAGGACGCCG	CCACCGCGCA	GACCTTGCAG	GCATTTCTGC	ACTGGGCGAT	6240
CACCGACGGC	AACAAGGCCT	CGTTCCTCGA	CCAGGTTCAT	TTCCAGCCGC	TGCCGCCCGC	6300
GGTGGTGAAG	TTGTCTGACG	CGTTGATCGC	GACGATTTCC	AGCGCTGAGA	TGAAGACCGA	6360
TGCCGCTACC	CTCGCGCAGG	AGGCAGGTAA	TTTCGAGCGG	ATCTCCGGCG	ACCTGAAAAC	6420
CCAGATCGAC	CAGGTGGAGT	CGACGGCAGG	TTCGTTGCAG	GGCCAGTGGC	GCGGCGCGGC	6480
GGGGACGGCC	GCCCAGGCCG	CGGTGGTGCG	CTTCCAAGAA	GCAGCCAATA	AGCAGAAGCA	6540
GGAACTCGAC	GAGATCTCGA	CGAATATTCG	TCAGGCCGGC	GTCCAATACT	CGAGGGCCGA	6600
CGAGGAGCAG	CAGCAGGCGC	TGTCCTCGCA	AATGGGCTTT	GTGCCCACAA	CGGCCGCCTC	6660

GCCGCCGTCG	ACCGCTGCAG	CGCCACCCGC	ACCGGCGACA	CCTGTTGCCC	CCCCACCACC	6720
GGCCGCCGCC	AACACGCCGA	ATGCCCAGCC	GGGCGATCCC	AACGCAGCAC	CTCCGCCGGC	6780
CGACCCGAAC	GCACCGCCGC	CACCTGTCAT	TGCCCCAAAC	GCACCCCAAC	CTGTCCGGAT	6840
CGACAACCCG	GTTGGAGGAT	TCAGCTTCGC	GCTGCCTGCT	GGCTGGGTGG	AGTCTGACGC	6900
CGCCCACTTC	GACTACGGTT	CAGCACTCCT	CAGCAAAACC	ACCGGGGACC	CGCCATTTCC	6960
CGGACAGCCG	CCGCCGGTGG	CCAATGACAC	CCGTATCGTG	CTCGGCCGGC	TAGACCAAAA	7020
GCTTTACGCC	AGCGCCGAAG	CCACCGACTC	CAAGGCCGCG	GCCCGGTTGG	GCTCGGACAT	7080
GGGTGAGTTC	TATATGCCCT	ACCCGGGCAC	CCGGATCAAC	CAGGAAACCG	TCTCGCTTGA	7140
CGCCAACGGG	GTGTCTGGAA	GCGCGTCGTA	TTACGAAGTC	AAGTTCAGCG	ATCCGAGTAA	7200
GCCGAACGGC	CAGATCTGGA	CGGGCGTAAT	CGGCTCGCCC	GCGGCGAACG	CACCGGACGC	7260
CGGGCCCCCT	CAGCGCTGGT	TTGTGGTATG	GCTCGGGACC	GCCAACAACC	CGGTGGACAA	7320
GGGCGCGGCC	AAGGCGCTGG	CCGAATCGAT	CCGGCCTTTG	GTCGCCCCGC	CGCCGGCGCC	7380
GGCACCGGCT	CCTGCAGAGC	CCGCTCCGGC	GCCGGCGCCG	GCCGGGGAAG	TCGCTCCTAC	7440
CCCGACGACA	CCGACACCGC	AGCGGACCTT	ACCGGCCTGA	GAATTCTGCA	GATATCCATC	7500
ACACTGGCGG	CCGCTCGAGC	ACCACCACCA	CCACCACTGA	GATCCGGCTG	CTAACAAAGC	7560
CCGAAAGGAA	GCTGAGTTGG	CTGCTGCCAC	CGCTGAGCAA	TAACTAGCAT	AACCCCTTGG	7620
GGCCTCTAAA	CGGGTCTTGA	GGGGTTTTTT	GCTGAAAGGA	GGAACTATAT	CCGGAT	7676

(2) INFORMATION FOR SEQ ID NO: 209:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 802 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:209:

Met Gly His His His His His Val Ile Asp Ile Ile Gly Thr Ser

Pro Thr Ser Trp Glu Gln Ala Ala Glu Ala Val Gln Arg Ala Arg 20 25 30

Asp	Ser	Val 35	Asp	Asp	Ile	Arg	Val 40	Ala	Arg	Val	Ile	Glu 45	Gln	Asp	Met
Ala	Val 50	Asp	Ser	Ala	Gly	Lys 55	Ile	Thr	Tyr	Arg	Ile 60	Lys	Leu	Glu	Val
Ser 65	Phe	Lys	Met	Arg	Pro 70	Ala	Gln	Pro	Arg	Gly 75	Ser	Lys	Pro	Pro	Ser 80
Gly	Ser	Pro	Glu	Thr 85	Gly	Ala	Gly	Ala	Gly 90	Thr	Val	Ala	Thr	Thr 95	Pro
Ala	Ser	Ser	Pro 100	Val	Thr	Leu	Ala	Glu 105	Thr	Gly	Ser		Leu 110	Leu	Tyr
Pro	Leu	Phe 115	Asn	Leu	Trp	Gly	Pro 120	Ala	Phe	His	Glu	Arg 125	Tyr	Pro	Asn
Val	Thr 130	Ile	Thr	Ala	Gln	Gly 135	Thr	Gly	Ser	Gly	Ala 140	Gly	Ile	Ala	Gln
Ala 145	Ala	Ala	Gly	Thr	Val 150	Asn	Ile	Gly	Ala	Ser 155	Asp	Ala	Tyr	Leu	Ser 160
Glu	Gly	Asp	Met	Ala 165	Ala	His	Lys	Gly	Leu 170	Met	Asn	Ile	Ala	Leu 175	Ala
Ile	Ser	Ala	Gln 180	Gln	Val	Asn	Tyr	Asn 185	Leu	Pro	Gly	Val	Ser 190	Glu	His
Leu	Lys	Leu 195	Asn	Gly	Lys	Val	Leu 200	Ala	Ala	Met	Tyr	Gln 205	Gly	Thr	Ile
Lys	Thr 210	Trp	Asp	Asp	Pro	Gln 215	Ile	Ala	Ala	Leu	Asn 220	Pro	Gly	Val	Asn
Leu 225	Pro	Gly	Thr	Ala	Val 230	Val	Pro	Leu	His	Arg 235		Asp	Gly	Ser	Gly 240
Asp	Thr	Phe	Leu	Phe 245	Thr	Gln	Tyr	Leu	Ser 250	Lys	Gln	Asp	Pro	Glu 255	Gly
Trp	Gly	Lys	Ser 260	Pro	Gly	Phe	Gly	Thr 265		Val	Asp	Phe	270	Ala	Val
Pro	Gly	Ala 275	Leu	Gly	Glu	Asn	Gly 280		Gly	Gly	Met	Val 285	Thr	Gly	Cys
Ala	Glu 290		Pro	Gly	Cys	Val 295		Туr	Ile	: Gly	300		Phe	. Leu	Asp
Gln 305		Ser	Gln	Arg	Gly 310	Leu	Gly	Glu	Ala	Gln 315		Gly	/ Asn	Ser	Ser 320
Glv	Asn	Phe	Leu	Leu	Pro	Asp	Ala	Gln	Ser	Ile	Glr	n Ala	a Ala	Alā	Ala

				325					330					335	
Gly	Phe	Ala	Ser 340	Lys	Thr	Pro	Ala	Asn 345	Gln	Ala	Ile	Ser	Met 350	Ile	Asp
Gly	Pro	Ala 355	Pro	Asp	Gly	Tyr	Pro 360	Ile	Ile	Asn	Tyr	Glu 365	Tyr	Ala	Ile
Val	Asn 370	Asn	Arg	Gln	Lys	Asp 375	Ala	Ala	Thr	Ala	Gln 380	Thr	Leu	Gln	Ala
Phe 385	Leu	His	Trp	Ala	Ile 390	Thr	Asp	Gly	Asn	Lys 395	Ala	Ser	Phe	Leu	Asp 400
Gln	Val	His	Phe	Gln 405	Pro	Leu	Pro	Pro	Ala 410	Val	Val	Lys	Leu	Ser 415	Asp
Ala	Leu	Ile	Ala 420	Thr	Ile	Ser	Ser	Ala 425	Glu	Met	Lys	Thr	Asp 430	Ala	Ala ,
Thr	Leu	Ala 435	Gln	Glu	Åla	Gly	Asn 440	Phe	Glu	Arg	Ile	Ser 445	Gly	Asp	Leu
Lys	Thr 450	Gln	Ile	Asp	Gln	Val 455	Glu	Ser	Thr	Ala	Gly 460	Ser	Leu	Gln	Gly
Gln 465	Trp	Arg	Gly	Ala	Ala 470	Gly	Thr	Ala	Ala	Gln 475	Ala	Ala	Val	Val	Arg 480
Phe	Gln	Glu	Ala	Ala 485	Asn	Lys	Gln	Lys	Gln 490	Glu	Leu	Asp	Glu	Ile 495	Ser
Thr	Asn	Ile	Arg 500	Gln	Ala	Gly	Val	Gln 505	Tyr	Ser	Arg	Ala	Asp 510	Glu	Glu
Gln	Gln	Gln 515	Ala	Leu	Ser	Ser	Gln 520	Met	Gly	Phe	Val	Pro 525	Thr	Thr	Ala
Ala	Ser 530	Pro	Pro	Ser	Thr	Ala 535	Ala	Ala	Pro	Pro	Ala 540	Pro	Ala	Thr	Pro
Val 545		Pro	Pro	Pro	Pro 550	Ala	Ala	Ala	Asn	Thr 555	Pro	Asr	n Ala	a Glr	Pro 560
Gly	Asp	Pro	Asn	Ala 565	Ala	Pro	Pro	Pro	Ala 570	Asp	Pro	Ası	n Alá	575	Pro
Pro	Pro	Val	Ile 580	Ala	Pro	Asn	Ala	Pro 585	Glr	n Pro	val	l Ar	g Ile 590	e Asp	Asn
Pro	Val	Gly 595	Gly	Phe	Ser	Phe	Ala 600		Pro	o Ala	a Gly	7 Tr	p Val	l Gl	ı Ser
Asp	Ala 610	Ala	His	Phe	Asp	Tyr 615		Ser	Ala	a Leu	1 Let 620	u Se O	r Ly	s Th	r Thr

Gly 625	Asp	Pro	Pro	Phe	Pro 630	Gly	Gln	Pro	Pro	Pro 635	Val	Ala	Asn	Asp	Thr 640
Arg	Ile	Val	Leu	Gly 645	Arg	Leu	Asp	Gln	Lys 650	Leu	Tyŕ	Ala	Ser	Ala 655	Glu
Ala	Thr	Asp	Ser 660	Lys	Ala	Ala	Ala	Arg 665	Leu	Gly	Ser	Asp	Met 670	Gly	Glu
Phe	Tyr	Met 675	Pro	Tyr	Pro	Gly	Thr 680	Arg	Ile	Asn	Gln	Glu 685	Thr	Val	Ser
Leu	Asp 690	Ala	Asn	Gly	Val	Ser 695	Gly	Ser	Ala	Ser	Tyr 700	Tyr	Glu	Val	Lys
Phe 705	Ser	Asp	Pro	Ser	Lys 710	Pro	Asn	Gly	Gln	Ile 715	Trp	Thr	Gly	Val	Ile 720
Gly	Ser	Pro	Ala	Ala 725	Asn	Ala	Pro	Asp	Ala 730	Gly	Pro	Pro	Gln	Arg 735	Trp
Phe	Val	Val	Trp 740	Leu	Gly	Thr	Ala	Asn 745	Asn	Pro	Val	Asp	Lys 750	Gly	Ala
Ala	Lys	Ala 755	Leu	Ala	Glu	Ser	Ile 760	Arg	Pro	Leu	Val	Ala 765	Pro	Pro	Pro
Ala	Pro 770	Ala	Pro	Ala	Pro	Ala 775	Glu	Pro	Ala	Pro	Ala 780	Pro	Ala	Pro	Ala
Gly 785	Glu	Val	Ala	Pro	Thr 790	Pro	Thr	Thr	Pro	Thr 795	Pro	Gln	Arg	Thr	Let 800
Pro	Ala														

CLAIMS

We claim:

- 1. A polypeptide comprising an antigenic portion of a soluble *M. tuberculosis* antigen, or a variant of said antigen that differs only in conservative substitutions and/or modifications, wherein said antigen has an N-terminal sequence selected from the group consisting of:
 - (a) Asp-Pro-Val-Asp-Ala-Val-Ile-Asn-Thr-Thr-Cys-Asn-Tyr-Gly-Gln-Val-Val-Ala-Ala-Leu (SEQ ID NO: 115);
 - (b) Ala-Val-Glu-Ser-Gly-Met-Leu-Ala-Leu-Gly-Thr-Pro-Ala-Pro-Ser (SEQ ID NO: 116);
 - (c) Ala-Ala-Met-Lys-Pro-Arg-Thr-Gly-Asp-Gly-Pro-Leu-Glu-Ala-Ala-Lys-Glu-Gly-Arg (SEQ ID NO: 17);
 - (d) Tyr-Tyr-Trp-Cys-Pro-Gly-Gln-Pro-Phe-Asp-Pro-Ala-Trp-Gly-Pro (SEQ ID NO: 118);
 - (e) Asp-Ile-Gly-Ser-Glu-Ser-Thr-Glu-Asp-Gln-Gln-Xaa-Ala-Val (SEQ ID NO: 119);
 - (f) Ala-Glu-Glu-Ser-Ile-Ser-Thr-Xaa-Glu-Xaa-Ile-Val-Pro (SEQ ID NO: 120);
 - (g) Asp-Pro-Glu-Pro-Ala-Pro-Pro-Val-Pro-Thr-Thr-Ala-Ala-Ser-Pro-Pro-Ser (SEQ ID NO: 121);
 - (h) Ala-Pro-Lys-Thr-Tyr-Xaa-Glu-Glu-Leu-Lys-Gly-Thr-Asp-Thr-Gly (SEQ ID NO: 122);
 - (i) Asp-Pro-Ala-Ser-Ala-Pro-Asp-Val-Pro-Thr-Ala-Ala-Gln-Leu-Thr-Ser-Leu-Leu-Asn-Ser-Leu-Ala-Asp-Pro-Asn-Val-Ser-Phe-Ala-Asn (SEQ ID NO: 123); and
 - (j) Ala-Pro-Glu-Ser-Gly-Ala-Gly-Leu-Gly-Gly-Thr-Val-Gln-Ala-Gly; (SEQ ID NO: 131)

wherein Xaa may be any amino acid.

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- 2. A polypeptide comprising an immunogenic portion of an *M. tuberculosis* antigen, or a variant of said antigen that differs only in conservative substitutions and/or modifications, wherein said antigen has an N-terminal sequence selected from the group consisting of:
 - (a) Asp-Pro-Pro-Asp-Pro-His-Gln-Xaa-Asp-Met-Thr-Lys-Gly-Tyr-Tyr-Pro-Gly-Gly-Arg-Arg-Xaa-Phe; (SEQ ID NO: 124) and
 - (b) Xaa-Tyr-Ile-Ala-Tyr-Xaa-Thr-Thr-Ala-Gly-Ile-Val-Pro-Gly-Lys-Ile-Asn-Val-His-Leu-Val; (SEQ ID NO: 132), wherein Xaa may be any amino acid.
- 3. A polypeptide comprising an antigenic portion of a soluble M. tuberculosis antigen, or a variant of said antigen that differs only in conservative substitutions and/or modifications, wherein said antigen comprises an amino acid sequence encoded by a DNA sequence selected from the group consisting of the sequences recited in SEQ ID NOS: 1, 2, 4-10, 13-25, 52, 94 and 96, the complements of said sequences, and DNA sequences that hybridize to a sequence recited in SEQ ID NOS: 1, 2, 4-10, 13-25, 52, 94 and 96 or a complement thereof under moderately stringent conditions.
- 4. A polypeptide comprising an antigenic portion of a *M. tuberculosis* antigen, or a variant of said antigen that differs only in conservative substitutions and/or modifications, wherein said antigen comprises an amino acid sequence encoded by a DNA sequence selected from the group consisting of the sequences recited in SEQ ID NOS: 26-51, 133, 134, 158-178 and 196, the complements of said sequences, and DNA sequences that hybridize to a sequence recited in SEQ ID NOS: 26-51, 133, 134, 158-178 and 196 or a complement thereof under moderately stringent conditions.
- 5. A DNA molecule comprising a nucleotide sequence encoding a polypeptide according to any one of claims 1-4.

- 6. A recombinant expression vector comprising a DNA molecule according to claim 5.
 - 7. A host cell transformed with an expression vector according to claim 6.
- 8. The host cell of claim 7 wherein the host cell is selected from the group consisting of *E. coli*, yeast and mammalian cells.
- 9. A method for detecting *M. tuberculosis* infection in a biological sample, comprising:
- (a) contacting a biological sample with one or more polypeptides according to any of claims 1-4; and
- (b) detecting in the sample the presence of antibodies that bind to at least one of the polypeptides, thereby detecting *M. tuberculosis* infection in the biological sample.
- 10. A method for detecting *M. tuberculosis* infection in a biological sample, comprising:
- (a) contacting a biological sample with a polypeptide having an N-terminal sequence selected from the group consisting of sequences provided in SEQ ID NO: 129 and 130; and
- (b) detecting in the sample the presence of antibodies that bind to at least one of the polypeptides, thereby detecting *M. tuberculosis* infection in the biological sample.
- 11. A method for detecting *M. tuberculosis* infection in a biological sample, comprising:
- (a) contacting a biological sample with one or more polypeptides encoded by a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198, the complements of said sequences, and DNA sequences that hybridize to a sequence recited in SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198; and

- (b) detecting in the sample the presence of antibodies that bind to at least one of the polypeptides, thereby detecting *M. tuberculosis* infection in the biological sample.
- 12. The method of any one of claims 9-11 wherein step (a) additionally comprises contacting the biological sample with a 38 kD *M. tuberculosis* antigen and step (b) additionally comprises detecting in the sample the presence of antibodies that bind to the 38 kD *M. tuberculosis* antigen.
- 13. The method of any one of claims 9-11 wherein the polypeptide(s) are bound to a solid support.
- 14. The method of claim 13 wherein the solid support comprises nitrocellulose, latex or a plastic material.
- 15. The method of any one of claims 9-11 wherein the biological sample is selected from the group consisting of whole blood, serum, plasma, saliva, cerebrospinal fluid and urine.
- 16. The method of claim 15 wherein the biological sample is whole blood or serum.
- 17. A method for detecting *M. tuberculosis* infection in a biological sample, comprising:
- (a) contacting the sample with at least two oligonucleotide primers in a polymerase chain reaction, wherein at least one of the oligonucleotide primers is specific for a DNA molecule according to claim 5; and
- (b) detecting in the sample a DNA sequence that amplifies in the presence of the oligonucleotide primers, thereby detecting *M. tuberculosis* infection.

- 18. The method of claim 17, wherein at least one of the oligonucleotide primers comprises at least about 10 contiguous nucleotides of a DNA molecule according to claim 5.
- 19. A method for detecting *M. tuberculosis* infection in a biological sample, comprising:
- (a) contacting the sample with at least two oligonucleotide primers in a polymerase chain reaction, wherein at least one of the oligonucleotide primers is specific for a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198; and
- (b) detecting in the sample a DNA sequence that amplifies in the presence of the first and second oligonucleotide primers, thereby detecting *M. tuberculosis* infection.
- 20. The method of claim 19, wherein at least one of the oligonucleotide primers comprises at least about 10 contiguous nucleotides of a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198.
- 21. The method of claims 17 or 19 wherein the biological sample is selected from the group consisting of whole blood, sputum, serum, plasma, saliva, cerebrospinal fluid and urine.
- 22. A method for detecting *M. tuberculosis* infection in a biological sample, comprising:
- (a) contacting the sample with one or more oligonucleotide probes specific for a DNA molecule according to claim 5; and
- (b) detecting in the sample a DNA sequence that hybridizes to the oligonucleotide probe, thereby detecting M. tuberculosis infection.

- 23. The method of claim 22 wherein the probe comprises at least about 15 contiguous nucleotides of a DNA molecule according to claim 5.
- 24. A method for detecting *M. tuberculosis* infection in a biological sample, comprising:
- (a) contacting the sample with one or more oligonucleotide probes specific for a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198; and
 - (b) detecting in the sample a DNA sequence that hybridizes to the oligonucleotide probe, thereby detecting M. tuberculosis infection.
- 25. The method of claim 24 wherein the oligonucleotide probe comprises at least about 15 contiguous nucleotides of a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198.
- 26. The method of claims 22 or 24 wherein the biological sample is selected from the group consisting of whole blood, sputum, serum, plasma, saliva, cerebrospinal fluid and urine.
 - 27. A method for detecting *M. tuberculosis* infection in a biological sample, comprising:
 - (a) contacting the biological sample with a binding agent which is capable of binding to a polypeptide according to any one of claims 1-4; and
 - (b) detecting in the sample a protein or polypeptide that binds to the binding agent, thereby detecting M. tuberculosis infection in the biological sample.
 - 28. A method for detecting *M. tuberculosis* infection in a biological sample, comprising:

- (a) contacting the biological sample with a binding agent which is capable of binding to a polypeptide having an N-terminal sequence selected from the group consisting of sequences provided in SEQ ID NO: 129 and 130; and
- (b) detecting in the sample a protein or polypeptide that binds to the binding agent, thereby detecting M. tuberculosis infection in the biological sample.
- 29. A method for detecting *M. tuberculosis* infection in a biological sample, comprising:
- (a) contacting the biological sample with a binding agent which is capable of binding to a polypeptide encoded by a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198, the complements of said sequences, and DNA sequences that hybridize to a sequence recited in SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198; and
- (b) detecting in the sample a protein or polypeptide that binds to the binding agent, thereby detecting M. tuberculosis infection in the biological sample.
- 30. The method of any one of claims 27-29 wherein the binding agent is a monoclonal antibody.
- 31. The method of any one of claims 27-29 wherein the binding agent is a polyclonal antibody.
 - 32. A diagnostic kit comprising:
 - (a) one or more polypeptides according to any of claims 1-4; and
 - (b) a detection reagent.
 - 33. A diagnostic kit comprising:
- (a) one or more polypeptides having an N-terminal sequence selected from the group consisting of sequences provided in SEQ ID NO: 129 and 130; and
 - (b) a detection reagent.

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- 34. A diagnostic kit comprising:
- (a) one or more polypeptides encoded by a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198, the complements of said sequences, and DNA sequences that hybridize to a sequence recited in SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198; and
 - (b) a detection reagent.
- 35. The kit of any one of claims 32-34 wherein the polypeptide(s) are immobilized on a solid support.
- 36. The kit of claim 35 wherein the solid support comprises nitrocellulose, latex or a plastic material.
- 37. The kit of any one of claims 32-34 wherein the detection reagent comprises a reporter group conjugated to a binding agent.
- 38. The kit of claim 37 wherein the binding agent is selected from the group consisting of anti-immunoglobulins, Protein G, Protein A and lectins.
- 39. The kit of claim 37 wherein the reporter group is selected from the group consisting of radioisotopes, fluorescent groups, luminescent groups, enzymes, biotin and dye particles.
- 40. A diagnostic kit comprising at least two oligonucleotide primers, at least one of the oligonucleotide primers being specific for a DNA molecule according to claim 5.

- 41. A diagnostic kit according to claim 40, wherein at least one of the oligonucleotide primers comprises at least about 10 contiguous nucleotide of a DNA molecule according to claim 5.
- 42. A diagnostic kit comprising a at least two oligonucleotide primers, at least one of the primers being specific for a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198.
- 43. A diagnostic kit according to claim 42, wherein at least one of the oligonucleotide primers comprises at least about 10 contiguous nucleotide of a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198.
- 44. A diagnostic kit comprising at least one oligonucleotide probe, the oligonucleotide probe being specific for a DNA molecule according to claim 5.
- 45. A kit according to claim 44, wherein the oligonucleotide probe comprises at least about 15 contiguous nucleotides of a DNA molecule according to claim 5.
- 46. A diagnostic kit comprising at least one oligonucleotide probe, the oligonucleotide probe being specific for a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198.
- 47. A kit according to claim 46, wherein the oligonucleotide probe comprises at least about 15 contiguous nucleotides of a DNA sequence selected from the group consisting of SEQ ID NOS: 3, 11, 12, 135, 136, 151-155, 184-188, 194-195 and 198.
- 48. A monoclonal antibody that binds to a polypeptide according to any of claims 1-4.

- 49. A polyclonal antibody that binds to a polypeptide according to any of claims 1-4.
- 50. A fusion protein comprising two or more polypeptides according to any one of claims 1-4.
- 51. A fusion protein comprising one or more polypeptides according to any one of claims 1-4 and ESAT-6 (SEQ ID NO: 99).
- 52. A fusion protein comprising a polypeptide having an N-terminal sequence selected from the group of sequences provided in SEQ ID NOS: 129 and 130.
- 53. A fusion protein comprising one or more polypeptides according to any one of claims 1-4 and the *M. tuberculosis* antigen 38 kD (SEQ ID NO: 150).
 - 54. A diagnostic kit comprising:
 - (a) one or more fusion proteins according to any one of claims 50-53; and
 - (b) a detection reagent.

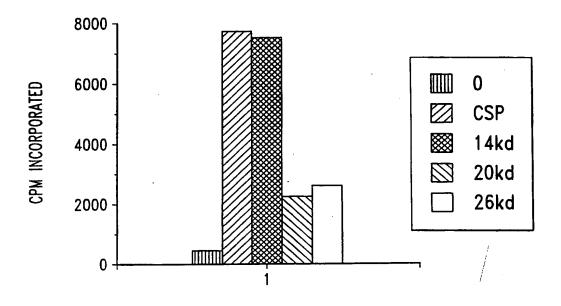


Fig. 1A-1

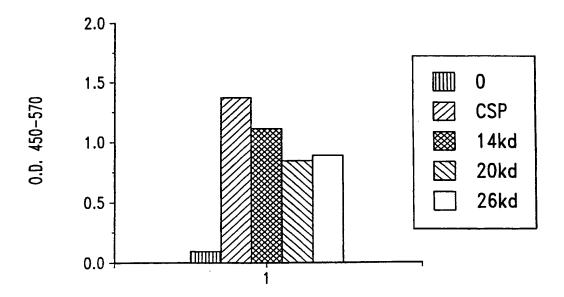


Fig. 1A-2

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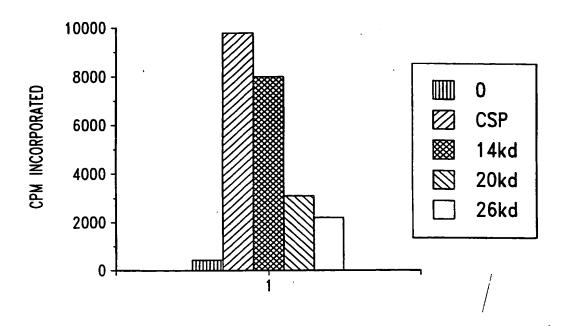


Fig. 1B-1

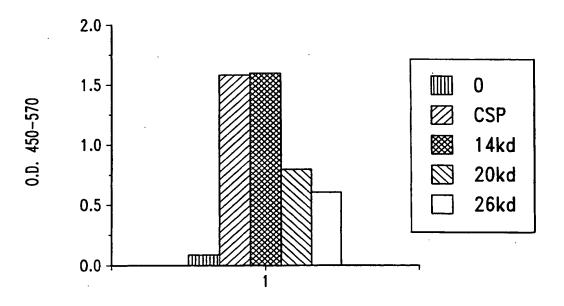
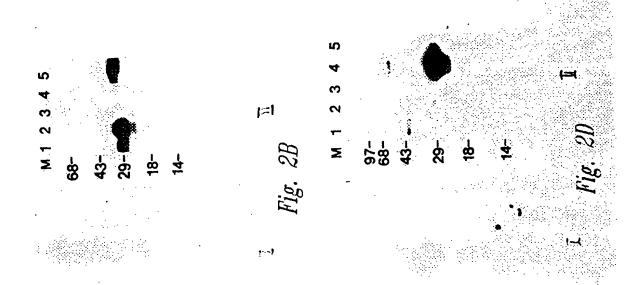
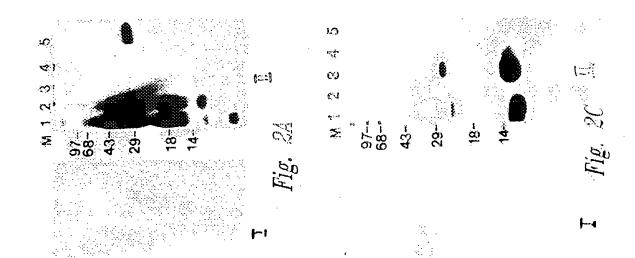
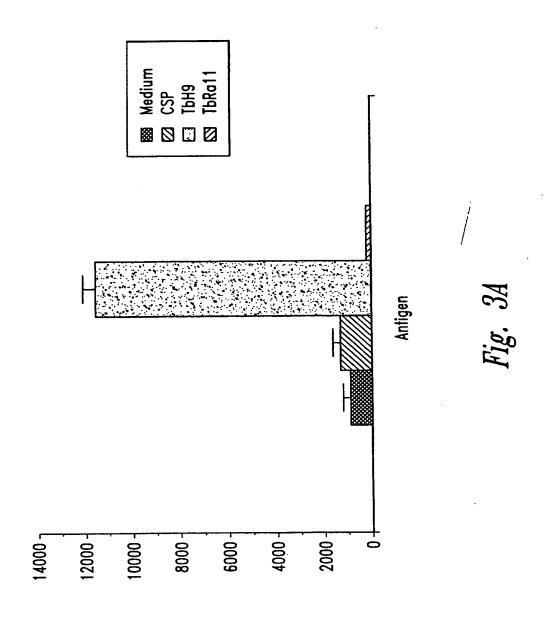


Fig. 1B-2

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CPM Incorporated

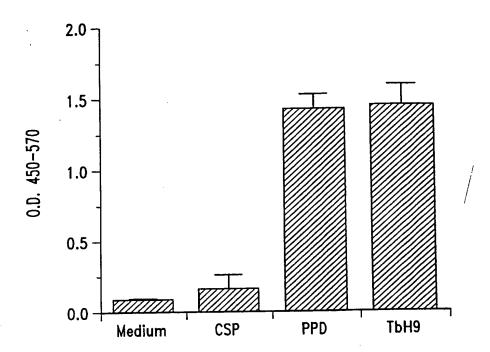
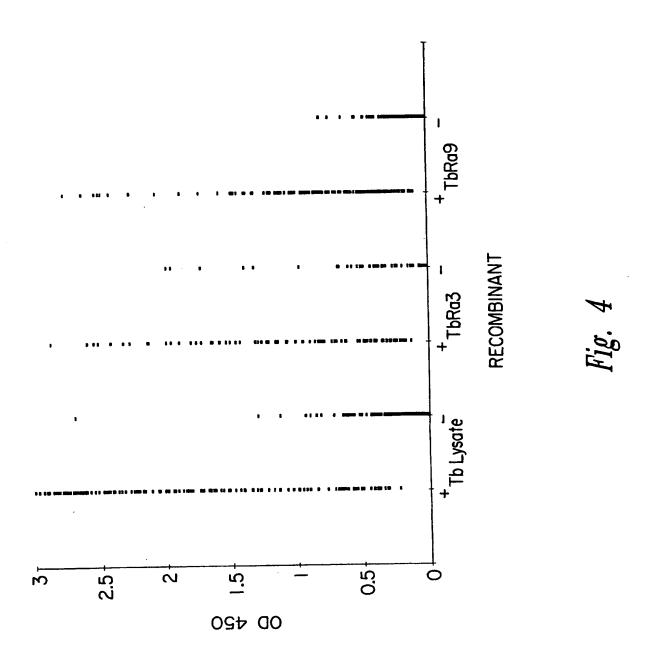
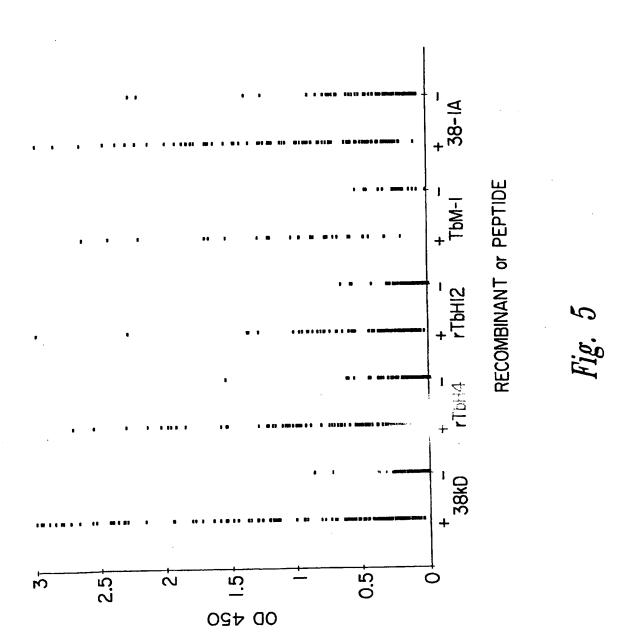
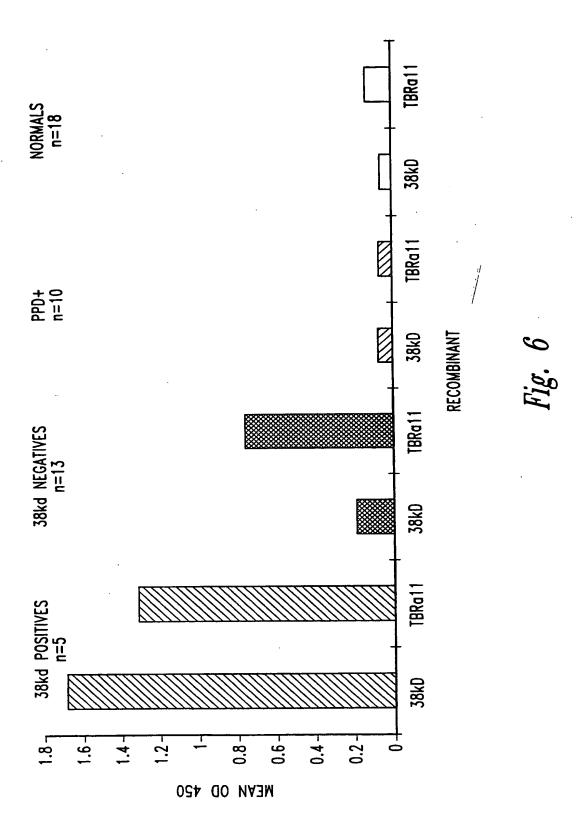


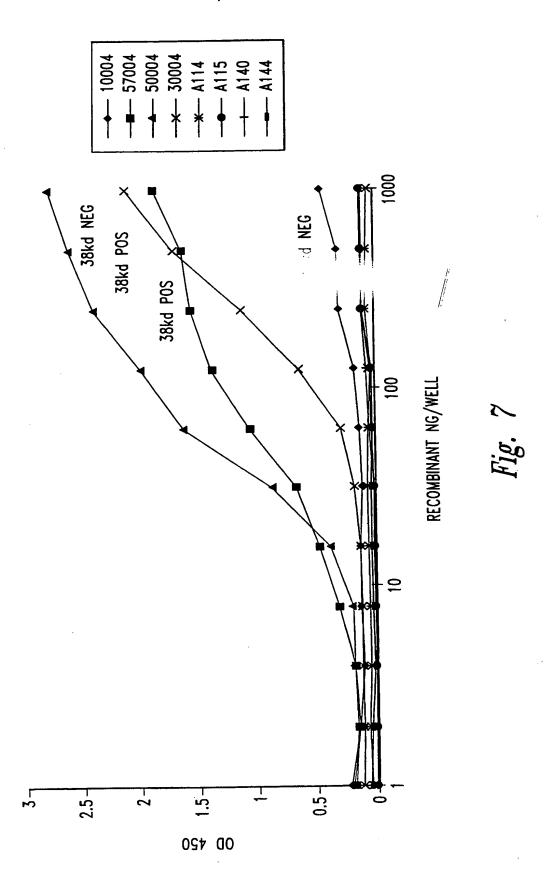
Fig. 3B





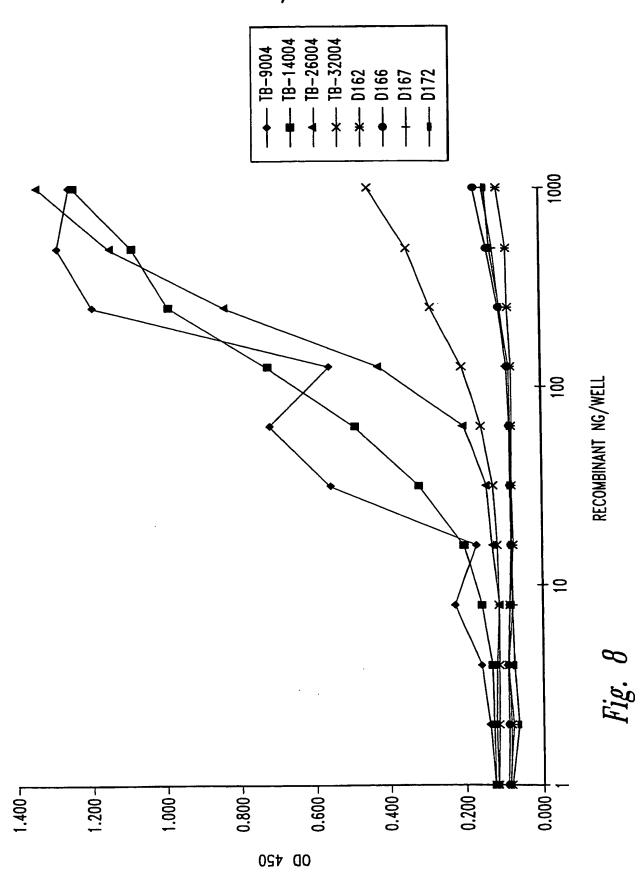


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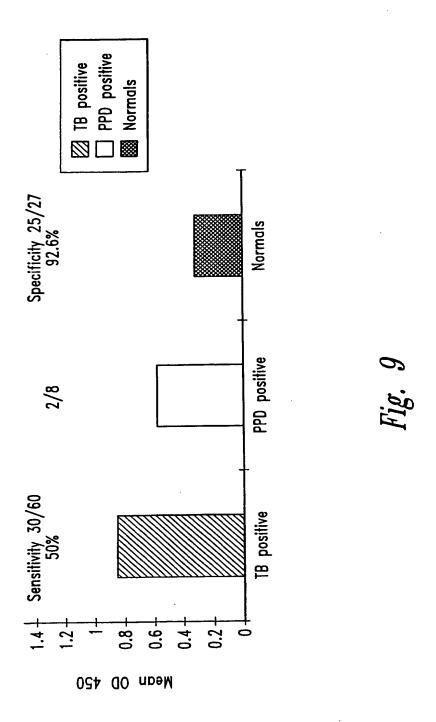


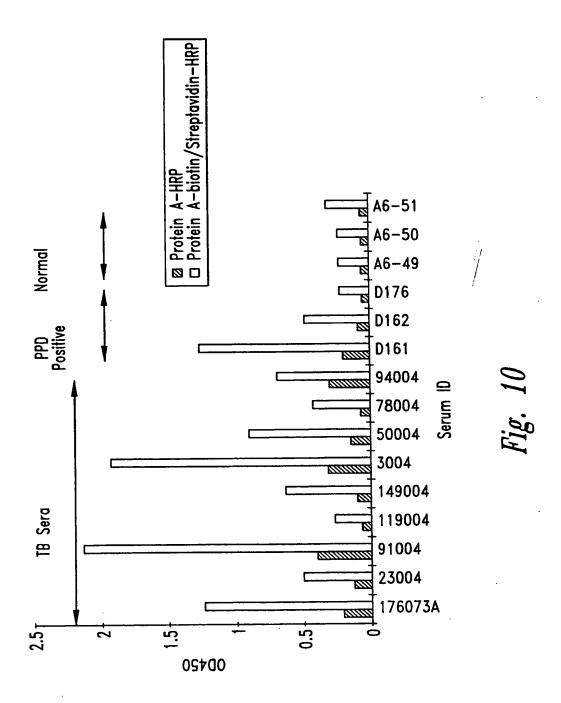
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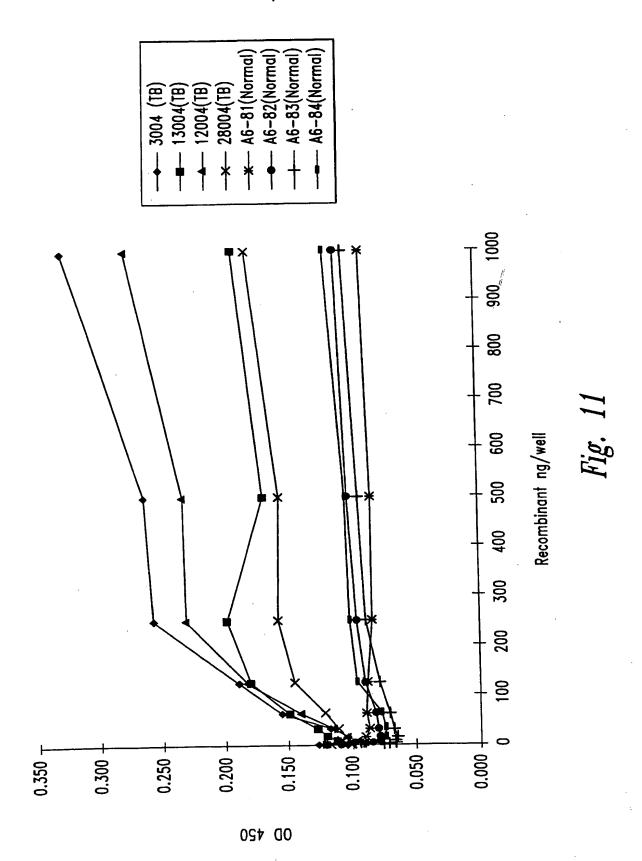
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(81) Designated States: AL, AM, AT, AU, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ARIPO patent (GH, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG).

(74) Agents: MAKI, David, J. et al.; Seed and Berry LLP,

6300 Columbia Center, 701 Fifth Avenue, Seattle, WA

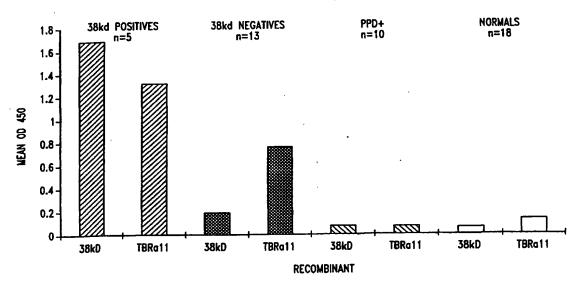
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(54) Title: COMPOUNDS AND METHODS FOR DIAGNOSIS OF TUBERCULOSIS



(57) Abstract

Compounds and methods for diagnosing tuberculosis are disclosed. The compounds provided include polypeptides that contain at least one antigenic portion of one or more M. tuberculosis proteins, and DNA sequences encoding such polypeptides. Diagnostic kits containing such polypeptides or DNA sequences and a suitable detection reagent may be used for the detection of M. tuberculosis infection in patients and biological samples. Antibodies directed against such polypeptides are also provided.

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Interna.. .al Application No PCT/US 97/18214

A. CLASSIFICATION OF SUBJECT MATTER IPC 6 C12N15/31 C07K14/35 C12N15/62 C12Q1/68 C07K16/12 G01N33/53 According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) C12N C07K C12Q G01N Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to claim No. Citation of document, with indication, where appropriate, of the relevant passages Category ° 1,3,5-9, EP 0 419 355 A (INNOGENETICS NV) 27 March Α 13-18, 21-23, 26,27, 30-32, 35-41, 44,45, 48,49 see abstract see page 24, line 45 - page 26, line 19 see page 56 - page 72; claims 50,51,54 WO 95 01441 A (STATENS SERUMSINSTITUT Α :ANDERSEN PETER (DK); ANDERSEN AASE BENGAAR) 12 January 1995 see abstract see page 20, line 13 - page 25, line 16 see page 73; claim 30 -/--Patent family members are listed in annex. Further documents are listed in the continuation of box C. Х X "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the Special categories of cited documents: *A* document defining the general state of the art which is not considered to be of particular relevance invention "E" earlier document but published on or after the international "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone filing date *L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such docucitation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or ments, such combination being obvious to a person skilled in the art. document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of mailing of the international search report Date of the actual completion of the international search 2 3, 06, 98 5 March 1998 Authorized officer Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo ni, Macchia, G Fax: (+31-70) 340-3016

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Internal ial Application No PCT/US 97/18214

C.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	WO 95 01440 A (STATENS SERUMINSTITUT ;HASLOEV KAARE (DK); ANDERSEN AASE BENGAARD) 12 January 1995	
A	ANDERSEN P. ET AL.: "Identification of immunodominant antigens during infection with Mycobacterium tuberculosis" SCANDINAVIAN JOURNAL OF IMMUNOLOGY, vol. 36, 1992, pages 823-831, XP002057751	
A	ANDERSEN A B ET AL: "STRUCTURE AND MAPPING OF ANTIGENIC DOMAINS OF PROTEIN ANTIGEN B, A 38,000-MOLECULAR-WEIGHT PROTEIN OF MYCOBACTERIUM TUBERCULOSIS" INFECTION AND IMMUNITY, vol. 57, no. 8, August 1989, pages 2481-2488, XP002026677 cited in the application see the whole document	12,53
A	WO 96 23885 A (PASTEUR INSTITUT ;LAQUEYRERIE ANNE (FR); MARCHAL GILLES (FR); PESC) 8 August 1996	
Α	WO 92 21758 A (PASTEUR INSTITUT) 10 December 1992	
A	AUSUBEL ET AL: "ISOLATION OF PROTEINS FOR MICROSEQUENCE ANALYSIS" CURRENT PROTOCOLS IN MOLECULAR BIOLOGY, 1993, pages 10.19.01-10.19.12, XP002026411 cited in the application	
A	YOUNG D B ET AL: "SCREENING OF A RECOMBINANT MYCOBACTERIAL DNA LIBRARY WITH POLYCLONAL ANTISERUM AND MOLECULAR WEIGHT ANALYSIS OF EXPRESSED ANTIGENS" INFECTION AND IMMUNITY, vol. 55, no. 6, June 1987, pages 1421-1425, XP002026410	
A	WO 94 00493 A (KAPOOR ARCHANA ;MUNSHI ANIL (US)) 6 January 1994	
A	FR 2 265 402 A (MITSUI PHARMACEUTICALS) 24 October 1975	·
A	FR 2 244 539 A (MITSUI PHARMACEUTICALS) 18 April 1975	

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C.(Continue Category °	ation) DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	ROMAIN ET AL: "PREPARATION OF TUBERCULIN ANTIGEN L" ANNALES DE L'INSTITUT PASTEUR / MICROBIOLOGIE, vol. 136B, 1985, pages 235-248, XP002026409	
P,X	WO 97 09429 A (CORIXA CORP) 13 March 1997	1,3,5-9, 12-18, 21-23, 26,27, 30-32, 35-41, 44,45, 48-51
	see abstract see page 173-181; claims	
P,X -	WO 97 09428 A (CORIXA CORP) 13 March 1997 see abstract see page 158 - page 163; claims	1,3,5-8

15

I national application No.

PCT/US 97/18214

B x I Observati ns where ertain claims were found unsearchabl (Continuation fitem 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see continuation-sheet
1. As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
A. X No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.: 1,3,5-9,12-18,21-23,26,27,30-32,35-41,44,45,48-51,53,54 all partially (subject 1. on next sheet)
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

1. Claims: 1, 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

A polypeptide comprising an antigenic portion of a soluble M. tuberculosis antigen or a variant, having an N-terminal aminoacid sequence as in Seq.ID:115 and/or encoded by a DNA molecule as in Seq.ID:96, complements of said sequence or sequences hybridizing to it. A DNA molecule comprising a sequence encoding said polypeptide. An expression vector comprising said DNA molecule, a host cell transformed with said expression vector. A method for detecting M. tuberculosis infection in a biological sample by detection of antibodies binding to said polypeptide or by detection of said polypeptide. A method for detecting M. tuberculosis infection in a biological sample by detection of said polypeptide. A method for detecting M. tuberculosis infection in a biological sample by detection of said DNA sequence. Diagnostic kits thereof. An antibody binding to said polypeptide. A fusion protein comprising said polypeptide. Diagnostic kit comprising said fusion protein.

2. Claims: 1, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:116.

3. Claims: 1, 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:(1)17 and 25.

4. Claims: 1, 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:118 and 24.

5. Claims: 1, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:119.

6. Claims: 1, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:120.

7. Claims: 1, 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:121 and 52.

8. Claims: 1, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:122.

9. Claims: 1, 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:123 and 94.

10. Claims: 1, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:131.

11. Claims: 2, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:124.

12. Claims: 2, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:132.

13. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:1.

14. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:2.

/.

15. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:4 and 17.

16. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:5.

17. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:6.

18. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:7.

19. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:8.

20. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:9.

21. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:10 and 13.

22. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:14.

23. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:15.

24. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:16.

25. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:18.

26. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:19.

27. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:20.

28. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:21.

29. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:22.

30. Claims: 3, 5-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:23.

31. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:26.

32. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:27.

33. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:28.

34. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:29.

35. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:30.

36. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:31.

37. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:32.

38. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:33.

39. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:34.

40. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:35.

41. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:36.

42. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:37.

43. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:38.

44. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:39.

45. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:40.

46. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID:41.

47. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:42.

48. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:43, 44 and 178.

49. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:45.

50. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID: 46.

51. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:47.

52. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:48.

53. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:49.

54. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:50.

55. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:51.

56. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:133.

57. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:134.

58. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:158.

59. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:159.

60. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:160.

61. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:161.

62. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:162.

63. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:163.

64. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID: 164 and 165.

65. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:166 and 167.

66. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:168 and 169.

67. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID: 170 and 171.

68. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq.ID:172 and 173.

69. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID: 174 and 175.

70. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID: 176 and 177.

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71. Claims: 4-9, 12-18, 21-23, 26, 27, 30-32, 35-41, 44, 45, 48-51, 53, 54 all partially.

Same as invention 1 but for Seq. ID: 196.

72. Claims: 10, 12-16, 28, 30, 31, 33, 35-39, 52, 54 all partially.

A method for detecting M. tuberculosis infection in a biological sample by detection of antibodies binding to a polypeptide having an N-terminal sequence as in Seq.ID:129, or by detection of a protein or polypeptide that binds to an agent binding to a polypeptide having an N-terminal sequence as in Seq.ID:129. Diagnostic kits thereof. A fusion protein comprising said polypeptide. Diagnostic kit comprising said fusion protein.

73. Claims: 10, 12-16, 28, 30, 31, 33, 35-39, 52, 54 all partially.

Same as invention 72 but for Seq.ID:130.

74. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

A method for detecting M. tuberculosis infection in a biological sample by detection of antibodies binding to a polypeptide encoded by a DNA sequence consisting of Seq.ID:3, complements or hybridizing sequences. A method for detecting M. tuberculosis infection in a biological sample by detection of said DNA sequence. A method for detecting M. tuberculosis infection in a biological sample by detection of a protein or polypeptide that binds to an agent binding to a polypeptide encoded by Seq.ID:3, complements or hybridizing sequences. Diagnostic kits thereof.

75. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:11.

76. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:12.

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77. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq. ID:135.

78. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:136.

79. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:151.

80. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:152.

81. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:153.

82. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:154 and 155.

83. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:184.

84. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:185.

85. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:186.

86. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:187.

87. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:188.

88. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:194 and 195.

89. Claims: 11-16, 19-21, 24-26, 29-31, 34-39, 42, 43, 46, 47 all partially.

Same as invention 74 but for Seq.ID:198.

Intermation on patent tamily members

Interna al Application No PCT/US 97/18214

Patent document cited in search report	Publication date	Patent family member(s)	Publication . date
EP 0419355 A	27-03-91	AU 6414390 A CA 2042016 A JP 9234096 A JP 2756368 B JP 4501811 T	18-04-91 20-03-91 09-09-97 25-05-98 02-04-92
WO 9501441 A	12-01-95	AU 682879 B AU 7068894 A CA 2165949 A EP 0706571 A NZ 267984 A	23-10-97 24-01-95 12-01-95 17-04-96 22-09-97
WO 9501440 A	12-01-95	AU 685133 B AU 7068694 A EP 0749486 A	15-01-98 24-01-95 27-12-96
WO 9623885 A	08-08-96	US 5714593 A AU 4667596 A CA 2210928 A EP 0807178 A	03-02-98 21-08-96 08-08-96 19-11-97
WO 9221758 A	10-12-92	FR 2677365 A CA 2110389 A EP 0589943 A JP 6508513 T	11-12-92 10-12-92 06-04-94 29-09-94
WO 9400493 A	06-01-94	US 5330754 A AU 689075 B AU 4651193 A EP 0649435 A JP 7508649 T US 5559011 A	19-07-94 26-03-98 24-01-94 26-04-95 28-09-95 24-09-96
FR 2265402 A	24-10-75	NONE	
FR 2244539 A	18-04-75	NONE	
WO 9709429 A	13-03-97	AU 7158796 A	27-03-97
WO 9709428 A	13-03-97	AU · 7158696 A	27-03-97

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